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### (57) Abstract

The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of Chlamydia pneumoniae, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, polypeptides encoded by such sequences, as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the Chlamydia pneumoniae genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing Chlamydia pneumoniae infection. The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular Chlamydia pneumoniae, infections.

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## CHLAMYDIA PNEUMONIAE GENOMIC SEQUENCE AND POLYPEPTIDES, FRAGMENTS THEREOF AND USES THEREOF, IN PARTICULAR FOR THE DIAGNOSIS, PREVENTION AND TREATMENT OF INFECTION

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The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of Chlamydia pneumoniae, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, 10 polypeptides encoded by such sequences, as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the Chlamydia pneumoniae genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing Chlamydia pneumoniae infection. 15 The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular Chlamydia pneumoniae, infections.

Comparative analysis of the sequence of the gene encoding the ribosomal 16S RNA has been widely used for the phylogenetic study of prokaryotes. This approach has made it possible to classify the Chlamydiae among the eubacteria, among which they represent a well-isolated group, with, nevertheless, a very weak link with the planctomyces. The Chlamydiae thus exhibit some unique characteristics within the eubacteria, in particular their development cycle and the structure of their 25 membranes. They have a unique two-phase cell cycle: the elementary body, a small extracellular form, attaches to the host and is phagocytosed; in the phagosome, it is converted to the replicative intracellular form, the reticulate body. The Chlamydiae are obligate intracellular bacteria which multiply in eukaryotic cells at the expense of their energy reserves and nucleotide pools; they are responsible for a wide variety of diseases in mammals and birds. The Chlamydiae are the only 30 members of the order Chlamydiales, of the family Chlamydiaceae and of the genus Chlamydia. Within the genus Chlamydia, four species are currently described: Chlamydia trachomatis, Chlamydia psittaci, Chlamydia pneumoniae and Chlamydia pecorum. These bacteria are grouped together and share biological and biochemical properties. Among them, only the first three infect humans, Chlamydia pecorum being a pathogen of ruminants.

The species Chlamydia psittaci infects many animals, in particular birds, and is transmissible to humans. It is responsible for atypical pneumonia, for hepatic and renal dysfunction, for endocarditis and for conjunctivitis.

The species Chlamydia trachomatis is the best characterized. Besides a murine strain, it is divided into two groups which are distinguishable by the nature of the diseases for which they are responsible: trachoma, genital attack and venereal lymphogranulomatosis. There are fifteen human serotypes of Chlamydia trachomatis (A, K) and LGV (L1, L2, L3). Strains A to C are mainly found in eye infections, whereas strains D to K and LGV are essentially responsible for genital entry infections. It should be mentioned that the LGV strains are responsible for systemic diseases. Historically, it was in 1906 that Halberstaeder and Von Provaseck discovered, in trachoma patients, the presence of inclusions in the cytoplasm of the cells derived from conjunctival scrapings. In 1940, Rake and Jones described these same inclusions in certain cells obtained by puncturing the ganglia from a patient suffering from venereal granulomatosis. Characterization of the Chlamydia trachomatis microorganism was only successfully carried out in 1957, after a series of isolations in cell cultures.

It was in 1983 that Chlamydia pneumoniae was recognized as a human pathogen (Grayston IT et al., 1986); since then, special attention has been paid to this bacterium and it is estimated (Gaydos CA et al., 1994) that 10% of pneumonias, and 5% of bronchitides and sinusites are attributable to Chlamydia pneumoniae (Aldous MB et al., 1992). More recently, the association of this bacterium with the pathogenesis of asthmatic disease and of cardiovascular impairments is increasingly of interest.

Serological studies have made it possible to observe that Chlamydia pneumoniae infection is common in children between 5 and 16 years of age. Before this age, it is rare to find antibodies; the increase in the number of individuals carrying antibodies is then correlated with age up to 20 years. Accordingly, 50% of adults are carriers of antibodies, it being possible for this prevalence to be as high as 75%. These figures are all the more striking since a first infection induces antibody levels of which the persistence over time is limited to 3 or at most 5 years, which suggests frequent reinfection during the entire lifespan. The annual seroconversion rate is about 8% between 8 and 12 years and about 6% between 12 and 16 years (Haidl et al., 1994). Before the age of 15 years, the seroprevalence of the disease is identical between both sexes. After this age, men are more frequently infected than women; this is true in all regions worldwide where such studies have been carried out.

These infections are geographically highly widespread, as shown by numerous studies carried out throughout the world (Kanamoto Y et al., 1991; Tong CY et al., 1993). Developed countries of the north such as Canada, Denmark and Norway have the lowest infection rates; conversely, the highest prevalence rates are found in the less developed countries of tropical regions where the infection may occur before the age of 5 years.

Humans are the only known reservoir for Chlamydia pneumoniae and it is probable that the infection is caused by direct transmission, respiratory secretions probably being responsible for this low-yield transmission (Aldous et al., 1992). The chain of transmission may also appear to be indirect (Kleemola M et al., 1988), suggesting that the infection is caused by an effective transmission, but also that asymptomatic carriers exist, which could explain the high prevalence of the disease.

Other studies (Mordhorst CH et al., 1992) show that the efficiency of the transmission varies according to the individuals and list cases of infection affecting all or the majority of members of one family or of a group of families. The period of incubation is several weeks, significantly longer in this regard than that of many other respiratory pathogenic agents. Although under conditions of high 5 relative humidity the infectivity of Chlamydia pneumoniae in the open air decreases rapidly, suggesting a direct mode of transmission under these conditions, it is probable that the transmission occurs in some cases indirectly since the microorganism can survive for up to 30 hours in a hostile environment (Falsey et al., 1993).

Clinical manifestations due to Chlamydia pneumoniae are essentially respiratory 10 diseases. Pneumonia and bronchitis are the most frequent because they are clinically patent: since etiological diagnosis is evoked in this case, the infectious agent is identified. The asymptomatic diseases are probably numerous (Grayston JT et al., 1992; Grayston JT et al., 1986; Thom DH et al., 1990). The disease then progresses via bronchitis or pneumonia; fever is absent at the time of examination but is sometimes reported by the patient. The degree of seriousness of the disease is 15 variable and in hospitalized patients, it is common to observe pleural effusion; a generalized infection may also be observed and, in severe cases, anatomicopathological examination shows Chlamydia pneumoniae diseases.

Other syndromes such as sinusitis (Hashiguchi K et al., 1992), purulent otitis media (Ogawa H et al., 1992), or pharyngitis (Huovinen P et al., 1989) have been described, as well as 20 infections with respiratory impairments similar to asthma (Hahn DL et al., 1991). Chlamydia pneumoniae has also been associated with sarcoidosis, with erythema nodosum (Sundelof et al., 1993) and one case of Guillain-Barré syndrome has even been described (Haidl et al., 1992). The involvement of Chlamydia pneumoniae in Reiter's syndrome has also been evaluated (Braun J et al., 1994).

The association of Chlamydia pneumoniae with coronary diseases and with myocardial infarction was first suspected from the observation of the high antibody level in 71% of patients having a heart disease (Shor A et al., 1992; Kuo CC et al., 1993; Puolakkainen M et al., 1993; Thomas GN et al., 1997). Studies carried out in several countries have shown similar results in patients with atheromatous impairments (Shor A et al., 1992; Kuo CC et al., 1993; Puolakkainen M 30 et al., 1993; Grayston JT et al., 1996; Casas-Ciria J et al., 1996; Thomas GN et al., 1997; Jackson LA et al., 1997) and in patients with carotid impairments. Anatomicopathological and microbiological studies have detected Chlamydia pneumoniae in the vessels. The electron microscope has made it possible to visualize the bacterium (Ladany S et al., 1989), which has in fact been demonstrated by other techniques such as PCR (Campbell LA et al., 1992; Kuo CC et al., 1993; Kuo CC et al., 1988). It 35 also appears that the bacterium is more frequently found in old atheromatous lesions. Other studies carried out on young subjects from 15 to 35 years have given the opportunity to study the coronary arteries of people without atherosclerosis, this observation not being possible in older subjects (the onset of the atheromatous disease is early). In these young subjects, the PCR studies did not find Chlamydia pneumoniae in subjects free of atheromatous disease, but revealed the presence of Chlamydia pneumoniae in two of the eleven subjects who showed early lesions and in six of the seven subjects who developed atheroma plaques. These studies therefore show that the atheroma plaque is very strongly correlated with the presence of Chlamydia pneumoniae, but the role played by the bacterium in vascular pathology is not yet defined.

The data relating to controlled clinical studies analysing the effect of treatments in Chlamydia pneumoniae infections are limited in number. Unlike penicillin, ampicillin or the sulphamides, erythromycin, tetracycline or doxycycline show an antibiotic activity in vitro against 10 Chlamydia pneumoniae. However, a treatment at high doses should be continued for several weeks in order to avoid a recurrence of the infection. Accordingly, the use of two new macrolides, clarithromycin and azithromycin, whose diffusion, bioavailability and half-life allow shorter and better tolerated cures, is nowadays preferred. In the absence of definitive proof based on the results of clinical studies, an effective, without recurrences, and well-tolerated treatment of Chlamydia pneumoniae infections therefore remains desirable.

An even more important need up until now relates to a specific and sensitive diagnosis, which can be carried out conveniently and rapidly, allowing early screening for the infection. Methods based on *Chlamydia pneumoniae* culture are slow and require a considerable know-how because of the difficulty involved in the collection, preservation and storage of the strain under appropriate conditions. Methods based on antigen detection (EIA, DFA) or on nucleic acid amplification (PCR) provide tests which are more suitable for laboratory practice. A reliable, sensitive and convenient test, which allows distinction between serogroups and a fortiori between *Chlamydia pneumoniae* species is therefore highly desirable.

This is all the more important since the symptoms of *Chlamydia pneumoniae* infection appear slowly, since all the pathologies associated with these infections have not yet been identified, and since, as has been mentioned above, an association is suspected between these infections and serious chronic infections, asthma or atherosclerosis.

No vaccine is yet available against *Chlamydia pneumoniae*: this is due to the labile nature of the antigens specific to the strain, which has so far prevented their specific identification.

Although the number of studies and of animal models developed is high, the antigens used have not induced sufficient protective immunity to lead to the development of human vaccines. In the case of *Chlamydia pneumoniae*, the role of the immune defense in the physiology and pathology of the disease should probably be understood in order to develop satisfactory vaccines.

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More detailed information relating to the biology of these strains, their interactions with their hosts, the associated phenomena of infectivity and those of escaping the immune defenses of the host in particular, and finally their involvement in the development of the these associated pathologies, will allow a better understanding of these mechanisms. In the light of the preceding text which shows

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in particular the limitations of the means of controlling *Chlamydia pneumoniae* infection, it is therefore at present essential, on the one hand, to develop molecular tools, in particular from a better genetic knowledge of *Chlamydia pneumoniae*, but also to develop new preventive and therapeutic treatments, new diagnostic methods and new vaccine strategies which are specific, effective and tolerated. This is precisely the object of the present invention.

The subject of the present invention is the nucleotide sequence having the sequence SEQ ID No. 1 of the *Chlamydia pneumoniae* genome. However, the invention is not limited to SEQ ID No. 1, but encompasses genomes and nucleotides encoding polypeptides of strain variants, polymorphisms, allelic variants, and mutants.

Thus, the subject of the present invention encompasses nucleotide sequences characterized in that they are chosen-from:

- a) the nucleotide sequence of SEQ ID No. 1, a nucleotide sequence exhibiting at least 99.9% identity with the sequence SEQ ID No. 1, the nucleotide sequence of the genomic DNA contained within ATCC Deposit No. \_\_\_\_, the nucleotide sequence of a clone insert within ATCC Deposit No. \_\_\_\_;
- b) a nucleotide sequence homologous to the sequence SEQ ID No. 1;
- c) a polynucleotide sequence that hybridizes to the nucleotide sequence of a) under conditions of high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are 20 as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65EC in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65EC, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 106 cpm of 32P-labeled probe. Alternatively, the hybridization step 25 can be performed at 65EC in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37EC for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50EC for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68EC for 15 minute intervals. Following 30 the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.
- 35 (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a

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temperature of 60EC in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50EC and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.

- d) a nucleotide sequence complementary to the sequence SEQ ID No. 1 or complementary to a nucleotide sequence as defined in a), b) or c) and a nucleotide sequence of their corresponding RNA;
- c) a nucleotide sequence of a representative fragment of the sequence SEQ ID No. 1, or of a representative fragment of the nucleotide sequence as defined in a), b), c) or d);
- f) a nucleotide sequence comprising a sequence as defined in a), b), c), d) or e);
- g) a nucleotide sequence capable of being obtained from a nucleotide sequence as defined in a), b), c), d), e) or f); and
- h) a modified nucleotide sequence of a nucleotide sequence as defined in a), b), c), d), e), f) or g).

Nucleotide sequence, polynucleotide or nucleic acid are understood to mean, according to the present invention, either a double-stranded DNA, a single-stranded DNA or products of transcription of the said DNAs.

It should be understood that the present invention does not relate to the genomic nucleotide sequences of *Chlamydia pneumoniae* taken in their natural environment, that is to say in the natural state. They are sequences which may have been isolated, purified or partially purified, by separation methods such as, for example, ion-exchange chromatography, molecular size exclusion chromatography or affinity chromatography, or alternatively fractionation techniques based on solubility in various solvents, or by genetic engineering methods such as amplification, cloning or subcloning, it being possible for the sequences of the invention to be carried by vectors.

The nucleotide sequence SEQ ID No. 1 was obtained by sequencing the *Chlamydia pneumoniae* genome by the method of directed sequencing after fluorescent automated sequencing of the inserts of clones and assembling of these sequences of nucleotide fragments (inserts) by means of softwares (cf. Examples). In spite of the high precision of the sequence SEQ ID No. 1, it is possible that it does not perfectly, 100% represent the nucleotide sequence of the *Chlamydia pneumoniae* genome and that a few rare sequencing errors or uncertainties still remain in the sequence SEQ ID No. 1. In the present invention, the presence of an uncertainty for an amino acid is designated by "Xaa" and that for a nucleotide is designated by "N" in the sequence listing below. These few rare errors or uncertainties could be easily detected and corrected by persons skilled in the art using the entire chromosome and/or its representative fragments according to the invention and standard

amplification, cloning and sequencing methods, it being possible for the sequences obtained to be easily compared, in particular by means of a computer software and using computer-readable media for recording the sequences according to the invention as described, for example, below. After correcting these possible rare errors or uncertainties, the corrected nucleotide sequence obtained would still exhibit at least 99.9% identity with the sequence SEQ ID No. 1. Such rare sequencing uncertainties are not present within the DNA contained within ATCC Deposit No. \_\_ or \_\_\_, and whatever rare sequence uncertainties that exist within SEQ ID No. 1 can routinely be corrected utilizing the DNA of the ATCC deposits.

Homologous nucleotide sequence for the purposes of the present invention is understood 10 to mean a nucleotide sequence having a percentage identity with the bases of the nucleotide sequence SEQ ID No. 1 of at least 80%, preferably 90% and 95%, this percentage being purely statistical and it being possible for the differences between the two nucleotide sequences to be distributed randomly and over their entire length. The said homologous sequences exhibiting a percentage identity with the bases of the nucleotide sequence SEQ ID No. 1 of at least 80%, preferably 90% and 95%, may 15 comprise, for example, the sequences corresponding to the genomic sequence or to the sequences of its representative fragments of a bacterium belonging to the Chlamydia family, including the species Chlamydia trachomatis, Chlamydia psittaci and Chlamydia pecorum mentioned above, as well as the sequences corresponding to the genomic sequence or to the sequences of its representative fragments of a bacterium belonging to the variants of the species Chlamydia pneumoniae. In the present 20 invention, the terms family and genus are mutually interchangeable, the terms variant, serotype, strain and subspecies are also mutually interchangeable. These homologous sequences may thus correspond to variations linked to mutations within the same species or between species and may correspond in particular to truncations, substitutions, deletions and/or additions of at least one nucleotide. The said homologous sequences may also correspond to variations linked to the degeneracy of the genetic code 25 or to a bias in the genetic code which is specific to the family, to the species or to the variant and which are likely to be present in Chlamydia.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8):2444-2448; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Altschul et al., 1993, Nature Genetics 3:266-272).

In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well known in the art (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997,

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Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the following task:

- (1)BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence database;
- (2)BLASTN compares a nucleotide query sequence against a nucleotide sequence database:
- (3)BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;
- (4)TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and
- (5)TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence 15 and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 20 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure, Washington: National Biomedical Research Foundation)

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a 25 high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268).

Nucleotide sequence complementary to a sequence of the invention is understood to mean any DNA whose nucleotides are complementary to those of the sequence of the invention, and whose orientation is reversed (antiparallel sequence).

The present invention further comprises fragments of the sequences of a) through f), above. Representative fragments of the sequences according to the invention will be understood to mean any nucleotide fragment having at least 8 successive nucleotides, preferably at least 12 successive nucleotides, and still more preferably at least 15 or at least 20 successive nucleotides of the sequence from which it is derived. It is understood that such fragments refer only to portions of SEQ 35 ID No. 1 that are not currently listed in a publicly available database.

Among these representative fragments, those capable of hybridizing under stringent conditions with a nucleotide sequence according to the invention are preferred. Hybridization under

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stringent conditions means that the temperature and ionic strength conditions are chosen such that they allow hybridization to be maintained between two complementary DNA fragments.

By way of illustration, high stringency conditions for the hybridization step for the purposes of defining the nucleotide fragments described above, are advantageously the following.

The hybridization is carried out at a preferred temperature of 65EC in the presence of SSC buffer, 1 × SSC corresponding to 0.15 M NaCl and 0.05 M Na citrate. The washing steps may be, for example, the following:

 $2 \times SSC$ , 0.1% SDS at room temperature followed by three washes with  $1 \times SSC$ , 0.1% SDS; 0.5 × SSC, 0.1% SDS; 0.1 × SSC, 0.1% SDS at 68EC for 15 minutes.

Intermediate stringency conditions, using, for example, a temperature of 60EC in the presence of a  $5 \times SSC$  buffer, or of low stringency, for example a temperature of 50EC in the presence of a  $5 \times SSC$  buffer, respectively require a lower overall complementarity for the hybridization between the two sequences.

The stringent hybridization conditions described above for a polynucleotide of about 300 bases in size will be adapted by persons skilled in the art for larger- or smaller-sized oligonucleotides, according to the teaching of Sambrook et al., 1989.

Among the representative fragments according to the invention, those which can be used as primer or probe in methods which make it possible to obtain homologous sequences or their representative fragments according to the invention, or to reconstitute a genomic fragment found to be incomplete in the sequence SEQ ID No. 1 or carrying an error or an uncertainty, are also preferred, these methods, such as the polymerase chain reaction (PCR), cloning and sequencing of nucleic acid being well known to persons skilled in the art. These homologous nucleotide sequences corresponding to mutations or to inter- or intra-species variations, as well as the complete genomic sequence or one of its representative fragments capable of being reconstituted, of course form part of the invention.

Among the said representative fragments, those which can be used as primer or probe in methods allowing diagnosis of the presence of *Chlamydia pneumoniae* or one of its associated microorganisms as defined below are also preferred.

The representative fragments capable of modulating, regulating, inhibiting or inducing the expression of a gene of *Chlamydia pneumoniae* or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia pneumoniae* or one of its associated microorganisms in the host cell and/or organism, are also preferred. Replication cycle is intended to designate invasion, multiplication, intracellular localization, in particular retention in the vacuole and inhibition of the process of fusion to the lysosome, and propagation of *Chlamydia pneumoniae* or one of its associated microorganisms from host cells to host cells.

Among the said representative fragments, those corresponding to nucleotide sequences corresponding to open reading frames, called ORF sequences (ORF for open reading frame), and

encoding polypeptides, such as for example, but without being limited thereto, the ORF sequences which will be later described, are finally preferred.

The representative fragments according to the invention may be obtained, for example, by specific amplification, such as PCR, or after digestion, with appropriate restriction enzymes, of nucleotide sequences according to the invention; these methods are in particular described in the manual by Sambrook et al., 1989. The said representative fragments may also be obtained by chemical synthesis when they are not too large in size and according to methods well known to persons skilled in the art. For example, such fragments can be obtained by isolating fragments of the genomic DNA of ATCC Deposit No. \_\_\_\_ or a clone insert present at this ATCC Deposit No. \_\_\_\_.

The representative fragments according to the invention may be used, for example, as primer, to reconstitute some of the said representative fragments, in-particular those in which a portion of the sequence is likely to be missing or imperfect, by methods well known to persons skilled in the art such as amplification, cloning or sequencing techniques.

Modified nucleotide sequence will be understood to mean any nucleotide sequence obtained by mutagenesis according to techniques well known to persons skilled in the art, and exhibiting modifications in relation to the normal sequences, for example mutations in the regulatory and/or promoter sequences for the expression of a polypeptide, in particular leading to a modification of the level of expression of the said polypeptide or to a modulation of the replicative cycle.

Modified nucleotide sequence will also be understood to mean any nucleotide sequence 20 encoding a modified polypeptide as defined below.

The subject of the present invention also includes *Chlamydia pneumoniae* nucleotide sequences characterized in that they are chosen from a nucleotide sequence of an open reading frame (ORF), that is, the ORF2 to ORF1297 sequences.

The ORF2 to ORF1297 nucleotide sequences are defined in Tables 1 and 2, *infra*, by their position on the sequence SEQ ID No. 1. For example, the ORF2 sequence is defined by the nucleotide sequence between the nucleotides at position 42 and 794 on the sequence SEQ ID No. 1, ends included. ORF2 to ORF1297 have been identified via homology analyses as well as via analyses of potential ORF start sites, as discussed in the examples below. It is to be understood that each identified ORF of the invention comprises a nucleotide sequence that spans the contiguous nucleotide sequence from the ORF stop codon immediately 3' to the stop codon of the preceding ORF and through the 5' codon to the next stop codon of SEQ ID No.:1 in-frame to the ORF nucleotide sequence. Table 2, *infra*, lists the beginning, end and potential start site of each of ORFs 1-1297. In one embodiment, the ORF comprises the contiguous nucleotide sequence spanning from the potential ORF start site downstream (that is, 3') to the ORF stop codon (or the ORF codon immediately adjacent to and upstream of the ORF stop codon). ORF2 to ORF1297 encode the polypeptides of SEQ ID No. 2 to SEQ ID No. 1291 and of SEQ ID No. 6844 to SEQ ID No. 6849, respectively.

Upon introduction of minor frameshifts, certain individual ORFs can comprise larger

"combined" ORFs. A list of such putative "combined" ORFs is shown in Table 3, below. For example, a combined ORF can comprise ORF 25, ORF 26 and ORF 27, including intervening inframe, nucleotide sequences. The order of ORFs (5' to 3'), within each "combined" ORF is as listed. It is to be understood that when ORF2 to ORF1297 are referred to herein, such reference is also meant to include "combined" ORFs. Polypeptide sequences encoded by such "combined" ORFs are also part of the present invention.

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Table 3
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ORF 25, ORF 26, ORF 27;

10 ORF 28, ORF 29, ORF 30;

ORF 31, ORF 32;

ORF 33, ORF 35;

ORF 466, ORF 467;

ORF 468, ORF 469;

15 ORF 477, ORF 476, ORF 474;

ORF 480, ORF 482;

ORF 483, ORF 485, ORF 486, ORF 500;

ORF 503, ORF 504, ORF 505;

ORF 506, ORF 507;

20 ORF 1211, ORF 647;

ORF 1286, ORF 1039;

ORF 691, ORF 690;

ORF 105, ORF 106;

ORF 170, ORF 171; ORF 394, ORF 393;

25 ORF 453, ORF 452, ORF 451;

ORF 526, ORF 525;

ORF 757, ORF 756, ORF 755;

ORF 856, ORF 855;

ORF 958, ORF 957;

30 ORF 915, ORF 914, ORF 913;

ORF 543, ORF 544;

ORF 1266, ORF 380;

ORF 745, ORF 744;

ORF 777, ORF 776;

35 ORF 343, ORF 1297, and representative fragments.

Table 1 also depicts the results of homology searches that compared the sequences of the

polypeptides encoded by each of the ORFs to sequences present in public published databases. It is understood that those polypeptides listed in Table 1 as exhibiting greater than about 95% identity to a polypeptide present in a publicly disclosed database are not considered part of the present invention; likewise in this embodiment, those nucleotide sequences encoding such polypeptides are not considered part of the invention. In another embodiment, it is understood that those polypeptides listed in Table 1 as exhibiting greater than about 99% identity to a polypeptide present in a publicly disclosed database are not considered part of the invention; likewise, in this embodiment, those nucleotide sequences encoding such polypeptides are not considered part of the invention.

The invention also relates to the nucleotide sequences characterized in that they comprise a nucleotide sequence chosen from:

- a) an ORF2 to ORF1297, a "combined" ORF nucleotide sequence, the nucleotide sequence of the genomic DNA contained within ATCC Deposit No. \_\_\_\_\_\_ or the nucleotide sequence of a clone insert in ATCC Deposit No. \_\_\_\_\_ according to the invention;
- b) a homologous nucleotide sequence exhibiting at least 80% identity across an entire ORF2 to ORF1297 nucleotide sequence according to the invention or as defined in a);
  - c) a polynucleotide sequence that hybridizes to ORF2 to ORF1297 under conditions of high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65EC in 20 buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65EC, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 10<sup>6</sup> cpm of <sup>32</sup>P-labeled probe. Alternatively, the hybridization step can be performed at 65EC in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 25 0.05 M Na citrate. Subsequently, filter washes can be done at 37EC for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50EC for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68EC for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high 30 stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1297. In one 35 embodiment, such sequences encode a Chlamydia pneumoniae polypeptide.
  - (ii) By way of example and not limitation, procedures using conditions of intermediate

stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60EC in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50EC and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1297. In one embodiment, such sequences encode a Chlamydia pneumoniae polypeptide.

- d) complementary or RNA nucleotide sequence corresponding to an ORF2 to ORF1297 sequence according to the invention or as defined in a), b) or c);
- e) a nucleotide sequence of a representative fragment of an ORF2 to ORF1297 sequence according to the invention or of a sequence as defined in a), b), c) or d);
- 15 f) a nucleotide sequence capable of being obtained from an ORF2 to ORF1297 sequence according to the invention or as defined in a), b), c), d) or e); and
  - g) a modified nucleotide sequence of an ORF2 to ORF1297 sequence according to the invention or as defined in a), b), c), d), e) or f);

As regards the homology with the ORF2 to ORF1297 nucleotide sequences, the homologous sequences exhibiting a percentage identity with the bases of one of the ORF2 to ORF1297 nucleotide sequences of at least 80%, preferably 90% and 95%, are preferred. Such homologous sequences are identified routinely via, for example, the algorithms described above and in the examples below. The said homologous sequences correspond to the homologous sequences as defined above and may comprise, for example, the sequences corresponding to the ORF sequences of a bacterium belonging to the Chlamydia family, including the species Chlamydia trachomatis, Chlamydia psittaci and Chlamydia pecorum mentioned above, as well as the sequences corresponding to the ORF sequences of a bacterium belonging to the variants of the species Chlamydia pneumoniae. These homologous sequences may likewise correspond to variations linked to mutations within the same species or between species and may correspond in particular to truncations, substitutions, deletions and/or additions of at least one nucleotide. The said homologous sequences may also correspond to variations linked to the degeneracy of the genetic code or to a bias in the genetic code which is specific to the family, to the species or to the variant and which are likely to be present in Chlamydia.

The invention comprises polypeptides encoded by a nucleotide sequence according to the invention, preferably by a representative fragment of the sequence SEQ ID No. 1 and corresponding to an ORF sequence, in particular the *Chlamydia pneumoniae* polypeptides, characterized in that they are chosen from the sequences SEQ ID No. 2 to SEQ ID No. 1291 or SEQ ID No. 6844 to SEQ ID No.

6849 and representative fragments thereof. However, the invention is not limited to polypeptides encoded by ORFs in SEQ ID No. 1 and its corresponding ORF sequences, but encompasses polypeptides of strain variants, polymorphisms, allelic variants, and mutants.

Thus, the invention also comprises the polypeptides characterized in that they comprise a polypeptide chosen from:

- a) a polypeptide encoded by a polynucleotide sequence in SEQ ID No. 1 (e.g., any polypeptide encoded by a polynucleotide sequence corresponding to ORF2 to ORF1297 and/or representative fragments thereof) according to the invention;
- b) a polypeptide homologous to a polypeptide according to the invention, or as defined in a);
- 10 c) a polypeptide encoded by a polynucleotide sequence that hybridizes to SEQ ID No. 1 or ORF2 to ORF1297 under high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65EC in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 15 0.02% BSA, and 500 μg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65EC, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 106 cpm of 32P-labeled probe. Alternatively, the hybridization step can be performed at 65EC in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37EC for 1 h in a solution containing 20 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50EC for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68EC for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, 25 Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably such polypeptide represents a homolog of a polypeptide encoded by ORF2 to ORF1297. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1297. In one embodiment, such 30 sequences encode a Chlamydia pneumoniae polypeptide.
- (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60EC in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50EC and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual,

Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1297. In one embodiment, such sequences encode a *Chlamydia pneumoniae* polypeptide.

- d) a fragment of at least 5 amino acids of a polypeptide according to the invention, or as defined in a), b) or c);
- e) a biologically active fragment of a polypeptide according to the invention, or as defined in a), b), c) or d); and
- 10 f) a modified polypeptide of a polypeptide according to the invention, as defined in a), b), c),d) or e).

In the present description, the terms polypeptide, peptide and protein are interchangeable.

It should be understood that the invention does not relate to the polypeptides in natural form, that is to say that they are not taken in their natural environment but that they may have been 15 isolated or obtained by purification from natural sources, or alternatively obtained by genetic recombination, or else by chemical synthesis and that they may, in this case, comprise nonnatural amino acids, as will be described below.

Homologous polypeptide will be understood to designate the polypeptides exhibiting, in relation to the natural polypeptide, certain modifications such as in particular a deletion, addition or substitution of at least one amino acid, a truncation, an extension, a chimeric fusion, and/or a mutation, or polypeptides exhibiting post-translational modifications. Among the homologous polypeptides, those whose amino acid sequence exhibits at least 80%, preferably 90%, homology or identity with the amino acid sequences of the polypeptides according to the invention are preferred. In the case of a substitution, one or more consecutive or nonconsecutive amino acids are replaced by "equivalent" amino acids. The expression "equivalent" amino acid is intended here to designate any amino acid capable of being substituted for one of the amino acids in the basic structure without, however, essentially modifying the biological activities of the corresponding peptides and as will be defined later.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8):2444-2448; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Altschul et al., 1993, Nature Genetics 3:266-272).

In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well know in the art (see,

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e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997, Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the following task:

- (1)BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence database;
- (2)BLASTN compares a nucleotide query sequence against a nucleotide sequence database;
- (3)BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;
- (4)TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and
- (5)TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.
- The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure, Washington: National Biomedical Research Foundation)

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268).

Equivalent amino acids may be determined either based on their structural homology 30 with the amino acids for which they are substituted, or on results of comparative tests of biological activity between the various polypeptides which may be carried out.

By way of example, there may be mentioned the possibilities of substitutions which may be carried out without resulting in a substantial modification of the biological activity of the corresponding modified polypeptides; the replacements, for example, of leucine with valine or isoleucine, of aspartic acid with glutamic acid, of glutamine with asparagine, of arginine with lysine, and the like, the reverse substitutions naturally being feasible under the same conditions.

The homologous polypeptides also correspond to the polypeptides encoded by the

homologous nucleotide sequences as defined above and thus comprise in the present definition the mutated polypeptides or polypeptides corresponding to inter- or intra-species variations which may exist in *Chlamydia*, and which correspond in particular to truncations, substitutions, deletions and/or additions of at least one amino acid residue.

Biologically active fragment of a polypeptide according to the invention will be understood to designate in particular a polypeptide fragment, as defined below, exhibiting at least one of the characteristics of the polypeptides according to the invention, in particular in that it is:

- capable of eliciting an immune response directed against Chlamydia pneumoniae; and/or

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- capable of being recognized by an antibody specific for a polypeptide according to the invention;
   and/or
- capable of binding to a polypeptide or to a nucleotide sequence of Chlamydia pneumoniae; and/or
- capable of modulating, regulating, inducing or inhibiting the expression of a gene of *Chlamydia* pneumoniae or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia pneumoniae* or one of its associated microorganisms in the host cell and/or organism; and/or
- capable of generally exerting an even partial physiological activity, such as for example a structural activity (cellular envelope, ribosome), an enzymatic (metabolic) activity, a transport activity, an activity in the secretion or in the virulence.

A polypeptide fragment according to the invention is understood to designate a 20 polypeptide comprising a minimum of 5 amino acids, preferably 10 amino acids or preferably 15 amino acids. It is to be understood that such fragments refer only to portions of polypeptides encoded by ORF2 to ORF1297 that are not currently listed in a publicly available database.

The polypeptide fragments according to the invention may correspond to isolated or purified fragments which are naturally present in *Chlamydia pneumoniae* or which are secreted by *Chlamydia pneumoniae*, or may correspond to fragments capable of being obtained by cleaving the said polypeptide with a proteolytic enzyme, such as trypsin or chymotrypsin or collagenase, or with a chemical reagent, such as cyanogen bromide (CNBr) or alternatively by placing the said polypeptide in a highly acidic environment, for example at pH 2.5. Such polypeptide fragments may be equally well prepared by chemical synthesis, using hosts transformed with an expression vector according to the invention containing a nucleic acid allowing the expression of the said fragments, placed under the control of appropriate elements for regulation and/or expression.

"Modified polypeptide" of a polypeptide according to the invention is understood to designate a polypeptide obtained by genetic recombination or by chemical synthesis as will be described below, exhibiting at least one modification in relation to the normal sequence. These modifications may in particular affect amino acids responsible for a specificity or for the efficiency of the activity, or responsible for the structural conformation, for the charge or for the hydrophobicity, and for the capacity for multimerization and for membrane insertion of the polypeptide according to

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the invention. It is thus possible to create polypeptides with an equivalent, an increased or a reduced activity, and with an equivalent, a narrower or a broader specificity. Among the modified polypeptides, there may be mentioned the polypeptides in which up to 5 amino acids may be modified, truncated at the N- or C-terminal end, or alternatively deleted, or else added.

As is indicated, the modifications of the polypeptide may have in particular the objective:

- of making it capable of modulating, regulating, inhibiting or inducing the expression of a gene of *Chlamydia*, in particular of *Chlamydia pneumoniae* and its variants, or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia*, in particular of *Chlamydia pneumoniae* and its variants, or one of its associated microorganisms, in the host cell and/or organism,
- of allowing its use in methods of biosynthesis or of biodegradation, or its incorporation into vaccine compositions,
- of modifying its bioavailability as a compound for therapeutic use.

The said modified polypeptides may also be used on any cell or microorganism for which the said modified polypeptides will be capable of modulating, regulating, inhibiting or inducing gene expression, or of modulating the growth or the replication cycle of the said cell or of the said microorganism. The methods allowing demonstration of the said modulations on eukaryotic or prokaryotic cells are well known to persons skilled in the art. The said cells or microorganisms will be chosen, in particular, from tumour cells or infectious microorganisms and the said modified polypeptides may be used for the prevention or treatment of pathologies linked to the presence of the said cells or of the said microorganisms. It is also clearly understood that the nucleotide sequences encoding the said modified polypeptides may be used for the said modulations, for example by the intermediacy of vectors according to the invention and which are described below, so as to prevent or to treat the said pathologies.

The above modified polypeptides may be obtained using combinatory chemistry, in which it is possible to systematically vary portions of the polypeptide before testing them on models, cell cultures or microorganisms for example, so as to select the compounds which are the most active or which exhibit the desired properties.

Chemical synthesis also has the advantage of being able to use:

- nonnatural amino acids, or
- nonpeptide bonds.

Accordingly, in order to extend the life of the polypeptides according to the invention, it may be advantageous to use nonnatural amino acids, for example in the D form, or alternatively amino acid analogues, in particular sulphur-containing forms for example.

Finally, the structure of the polypeptides according to the invention, its homologous or modified forms, as well as the corresponding fragments may be integrated into chemical structures of the polypeptide type and the like. Accordingly, it may be advantageous to provide at the N- and C-

terminal ends compounds which are not recognized by proteases.

Also forming part of the invention are the nucleotide sequences encoding a polypeptide according to the invention. Described below are ORF nucleotide sequences encoding polypeptides exhibiting particularly preferable characteristics. For each group of preferred ORFS described below, 5 it is to be understood that in addition to the individual ORFs listed, in instances wherein such ORFS are present as part of "combined" ORFs, the "combined" ORFs are also to be included within the preferred group.

More particularly, the subject of the invention is nucleotide sequences, characterized in that they encode a polypeptide of the cellular envelope, preferably of the outer cellular envelope of 10 Chlamydia pneumoniae or one of its representative fragments, such as for example the predominant proteins of the outer membrane, the adhesion proteins or the proteins entering into the composition of the Chlamydia wall. Among these sequences, the sequences comprising a nucleotide sequence chosen from the following sequences are most preferred:

ORF15; ORF25; ORF26; ORF27; ORF28; ORF29; ORF30; ORF31; ORF32; ORF33; ORF35; 15 ORF68; ORF124; ORF275; ORF291; ORF294; ORF327; ORF342; ORF364; ORF374; ORF380; ORF414; ORF439; ORF466; ORF467; ORF468; ORF469; ORF470; ORF472; ORF474; ORF476; ORF477; ORF478; ORF479; ORF480; ORF482; ORF485; ORF500; ORF501; ORF503; ORF504; ORF505; ORF506; ORF520; ORF578; ORF580; ORF581; ORF595; ORF596; ORF597; ORF737; ORF830; ORF834; ORF836; ORF893; ORF917; ORF932; ORF976; ORF1035; ORF1045; ORF1090 20 and one of their representative fragments.

The structure of the cytoplasmic membranes and of the wall of bacteria is dependent on the associated proteins. The structure of the cytoplasmic membrane makes it impermeable to water, to water-soluble substances and to small-sized molecules (ions, small inorganic molecules, peptides or proteins). To enter into or to interfere with a cell or a bacterium, a ligand must establish a special 25 relationship with a protein anchored in the cytoplasmic membrane (the receptor). These proteins which are anchored on the membrane play an important role in metabolism since they control the exchanges in the bacterium. These exchanges apply to molecules of interest for the bacterium (small molecules such as sugars and small peptides) as well as undesirable molecules for the bacterium such as antibiotics or heavy metals.

The double lipid layer structure of the membrane requires the proteins which are inserted therein to have hydrophobic domains of about twenty amino acids forming an alpha helix. Predominantly hydrophobic and potentially transmembrane regions may be predicted from the primary sequence of the proteins, itself deduced from the nucleotide sequence. The presence of one or more putative transmembrane domains raises the possibility for a protein to be associated with the 35 cytoplasmic membrane and to be able to play an important metabolic role therein or alternatively for the protein thus exposed to be able to exhibit potentially protective epitopes.

If the proteins inserted into the membrane exhibit several transmembrane domains

capable of interacting with one another via electrostatic bonds, it then becomes possible for these proteins to form pores which go across the membrane which becomes permeable for a number of substances. It should be noted that proteins which do not have transmembrane domains may also be anchored by the intermediacy of fatty acids in the cytoplasmic membrane, it being possible for the breaking of the bond between the protein and its anchor in some cases to be responsible for the release of the peptide outside the bacterium.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* transmembrane polypeptide or one of its representative fragments, having between 1 and 3 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

10 nucleotide sequence chosen from the following sequences: ORF2; ORF3; ORF6; ORF9; ORF10; ORF11; ORF13; ORF14; ORF16; ORF18; ORF19; ORF20; ORF21; ORF22; ORF25; ORF27; ORF28; ORF29; ORF30; ORF31; ORF32; ORF33; ORF34; ORF35; ORF37; ORF39; ORF41; ORF42; ORF44; ORF45; ORF46; ORF47; ORF48; ORF49; ORF50; ORF53; ORF54; ORF56; ORF57; ORF59; ORF60; ORF61; ORF62; ORF63; ORF64; 15 ORF65; ORF66; ORF69;; ORF72; ORF73; ORF74; ORF76; ORF77; ORF78; ORF79; ORF80; ORF82; ORF84; ORF85; ORF86; ORF88; ORF89; ORF90; ORF91; ORF92; ORF93; ORF95; ORF96; ORF98; ORF99; ORF100; ORF101; ORF102; ORF103; ORF104; ORF105; ORF106; ORF107; ORF108; ORF114; ORF117; ORF118; ORF122; ORF123; ORF124; ORF125; ORF129; ORF130; ORF131; ORF132; ORF133; ORF134; ORF135; ORF137; ORF138; ORF139; ORF140; 20 ORF141; ORF142; ORF143; ORF145; ORF146; ORF147; ORF150; ORF151; ORF152; ORF156; ORF157; ORF158; ORF159; ORF160; ORF161; ORF162; ORF164; ORF166; ORF167; ORF170; ORF173; ORF175; ORF176; ORF178; ORF179; ORF180; ORF182; ORF183; ORF184; ORF185; ORF186; ORF187; ORF188; ORF189; ORF190; ORF191; ORF192; ORF194; ORF195; ORF196; ORF197; ORF198; ORF199; ORF200; ORF201; ORF202; ORF205; ORF207; ORF208; ORF209; 25 ORF210; ORF212; ORF215; ORF219; ORF220; ORF224; ORF226; ORF227; ORF228; ORF231; ORF232; ORF233; ORF234; ORF235; ORF236; ORF238; ORF239; ORF240; ORF241; ORF242; ORF244; ORF247; ORF251; ORF252; ORF253; ORF255; ORF256; ORF257; ORF258; ORF260; ORF262; ORF263; ORF266; ORF267; ORF268; ORF269; ORF270; ORF273; ORF274; ORF276; ORF278; ORF279; ORF280; ORF281; ORF282; ORF283; ORF284; ORF286; ORF287; ORF289; 30 ORF290; ORF291; ORF293; ORF294; ORF297; ORF304; ORF305; ORF307; ORF308; ORF309; ORF310; ORF311; ORF313; ORF314; ORF315; ORF316; ORF318; ORF319; ORF320; ORF321; ORF322: ORF323: ORF324; ORF325; ORF326; ORF331; ORF332; ORF336; ORF338; ORF339; ORF341; ORF344; ORF345; ORF346; ORF350; ORF352; ORF353; ORF356; ORF357; ORF358; ORF359; ORF360; ORF362; ORF365; ORF366; ORF367; ORF370; ORF372; ORF373; ORF376; 35 ORF377; ORF378; ORF379; ORF381; ORF382; ORF383; ORF384; ORF385; ORF386; ORF387; ORF390; ORF392; ORF393; ORF394; ORF396; ORF398; ORF399; ORF400; ORF404; ORF408; ORF410; ORF411; ORF413; ORF416; ORF417; ORF418; ORF420; ORF422; ORF424; ORF427;

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ORF428; ORF429; ORF430; ORF431; ORF433; ORF434; ORF437; ORF440; ORF441; ORF442:
   ORF443; ORF444; ORF445; ORF447; ORF450; ORF451; ORF452; ORF455; ORF456; ORF459;
   ORF460; ORF461; ORF462; ORF463; ORF464; ORF465; ORF467; ORF469; ORF471; ORF474;
   ORF475; ORF476; ORF477; ORF479; ORF482; ORF483; ORF484; ORF485; ORF486; ORF487;
 5 ORF488; ORF491; ORF493; ORF494; ORF497; ORF498; ORF499; ORF503; ORF508; ORF509;
   ORF510: ORF512; ORF514; ORF515; ORF516; ORF517; ORF518; ORF520; ORF521; ORF523;
   ORF525: ORF527; ORF528; ORF529; ORF530; ORF531; ORF533; ORF534; ORF535; ORF536;
   ORF537; ORF540; ORF541; ORF543; ORF544; ORF545; ORF546; ORF548; ORF549; ORF551;
   ORF553; ORF554; ORF555; ORF556; ORF557; ORF558; ORF559; ORF560; ORF562; ORF563;
10 ORF564; ORF565; ORF566; ORF569; ORF571; ORF573; ORF576; ORF577; ORF581; ORF583;
   ORF584: ORF585; ORF586; ORF588; ORF591; ORF592; ORF594, ORF595; ORF596; ORF597;
   ORF599; ORF600; ORF603; ORF605; ORF608; ORF614; ORF615; ORF620; ORF621; ORF622;
   ORF623; ORF624; ORF625; ORF629; ORF630; ORF631; ORF633; ORF634; ORF637; ORF642;
   ORF644; ORF645; ORF647; ORF648; ORF652; ORF654; ORF655; ORF657; ORF658; ORF659;
15 ORF660: ORF661: ORF664: ORF665; ORF666; ORF667; ORF670; ORF671; ORF672; ORF673;
   ORF674; ORF676; ORF679; ORF681; ORF684; ORF687; ORF688; ORF689; ORF690; ORF693;
   ORF694: ORF695: ORF696: ORF697: ORF698: ORF699; ORF700; ORF701; ORF703; ORF705;
   ORF706; ORF707; ORF708; ORF710; ORF712; ORF715; ORF716; ORF717; ORF718; ORF719;
   ORF721; ORF722; ORF723; ORF725; ORF726; ORF727; ORF728; ORF729; ORF730; ORF731;
20 ORF733; ORF736; ORF737; ORF738; ORF740; ORF741; ORF742; ORF743; ORF747; ORF748;
    ORF750; ORF752; ORF754; ORF755; ORF756; ORF757; ORF759; ORF760; ORF761; ORF762;
    ORF763; ORF764; ORF765; ORF766; ORF767; ORF768; ORF772; ORF774; ORF775; ORF777;
    ORF781: ORF783: ORF788: ORF791; ORF792; ORF793; ORF794; ORF795; ORF796; ORF797;
    ORF798; ORF802; ORF803; ORF806; ORF807; ORF808; ORF809; ORF810; ORF811;
25 ORF813; ORF814; ORF815; ORF816; ORF817; ORF819; ORF820; ORF821; ORF823; ORF824;
    ORF827; ORF829; ORF830; ORF831; ORF833; ORF834; ORF835; ORF837; ORF844; ORF845;
    ORF846: ORF847; ORF848; ORF849; ORF850; ORF851; ORF852; ORF854; ORF855; ORF856;
    ORF857; ORF859; ORF860; ORF862; ORF865; ORF866; ORF868; ORF869; ORF870; ORF871;
    ORF872; ORF874; ORF877; ORF878; ORF879; ORF880; ORF881; ORF882; ORF884; ORF885;
30 ORF888; ORF889; ORF890; ORF891; ORF892; ORF894; ORF895; ORF896; ORF897; ORF899;
    ORF900; ORF902; ORF903; ORF904; ORF905; ORF909; ORF910; ORF912; ORF913; ORF914;
    ORF915: ORF917: ORF918: ORF919: ORF921: ORF923; ORF924; ORF926; ORF927; ORF928;
    ORF929: ORF930: ORF931: ORF937: ORF938: ORF939: ORF941; ORF943; ORF948; ORF951;
    ORF952; ORF953; ORF958; ORF960; ORF963; ORF964; ORF965; ORF968; ORF970; ORF974;
35 ORF975; ORF977; ORF979; ORF980; ORF981; ORF983; ORF984; ORF985; ORF987; ORF989;
    ORF992; ORF993; ORF997; ORF998; ORF999; ORF1001; ORF1002; ORF1004; ORF1005;
    ORF1009; ORF1013; ORF1014; ORF1015; ORF1016; ORF1019; ORF1021; ORF1023; ORF1024;
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ORF1029; ORF1031; ORF1033; ORF1034; ORF1039; ORF1041; ORF1042; ORF1045; ORF1047; ORF1049; ORF1051; ORF1052; ORF1053; ORF1054; ORF1056; ORF1059; ORF1061; ORF1062; ORF1063; ORF1064; ORF1065; ORF1067; ORF1075; ORF1077; ORF1078; ORF1079; ORF1080; ORF1081; ORF1089; ORF1095; ORF1097; ORF1098; ORF1099; ORF1101; ORF1102; ORF1103; ORF1106; ORF1107; ORF1108; ORF1109; ORF1110; ORF1113; ORF1116; ORF1118; ORF1119; ORF1121; ORF1123; ORF1124; ORF1126; ORF1128; ORF1130; ORF1131; ORF1133; ORF1134; ORF1136; ORF1137 and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a Chlamydia pneumoniae transmembrane polypeptide or one of its 10 representative fragments, having between 4 and 6 transmembrane domains and in that they comprise a nucleotide sequence chosen-from the following sequences: ORF5; ORF7; ORF8; ORF15; ORF36; ORF38; ORF51; ORF55; ORF58; ORF67; ORF70; ORF81; ORF97; ORF110; ORF111; ORF115; ORF119; ORF126; ORF128; ORF148; ORF155; ORF163; ORF165; ORF168; ORF169; ORF171; ORF172; ORF174; ORF177; ORF181; ORF193; ORF203; 15 ORF213; ORF214; ORF216; ORF217; ORF221; ORF222; ORF225; ORF229; ORF243; ORF246; ORF248; ORF254; ORF261; ORF285; ORF288; ORF292; ORF296; ORF298; ORF299; ORF301; ORF303; ORF317; ORF328; ORF329; ORF351; ORF354; ORF355; ORF364; ORF371; ORF374; ORF375; ORF391; ORF395; ORF401; ORF403; ORF405; ORF409; ORF414; ORF419; ORF421; ORF423; ORF425; ORF438; ORF448; ORF453; ORF458; ORF466; ORF468; ORF470; ORF480; 20 ORF489; ORF490; ORF496; ORF501; ORF504; ORF505; ORF506; ORF511; ORF513; ORF519; ORF526; ORF532; ORF538; ORF539; ORF547; ORF550; ORF561; ORF568; ORF570; ORF574; ORF578; ORF579; ORF580; ORF582; ORF589; ORF593; ORF598; ORF601; ORF604; ORF610; ORF613; ORF617; ORF626; ORF632; ORF635; ORF638; ORF640; ORF641; ORF646; ORF649; ORF650; ORF651; ORF686; ORF711; ORF724; ORF732; ORF734; ORF744; ORF745; ORF749; 25 ORF751; ORF769; ORF770; ORF771; ORF773; ORF776; ORF779; ORF780; ORF785; ORF787; ORF789; ORF801; ORF805; ORF812; ORF822; ORF825; ORF826; ORF839; ORF841; ORF843; ORF853; ORF861; ORF875; ORF876; ORF886; ORF893; ORF898; ORF906; ORF907; ORF908; ORF920; ORF922; ORF925; ORF933; ORF935; ORF936; ORF944; ORF946; ORF947; ORF954; ORF959; ORF961; ORF966; ORF967; ORF972; ORF978; ORF995; ORF996; ORF1000; ORF1003; 30 ORF1010; ORF1011; ORF1012; ORF1017; ORF1020; ORF1030; ORF1036; ORF1038; ORF1043; ORF1046; ORF1048; ORF1050; ORF1058; ORF1071; ORF1073; ORF1084; ORF1085; ORF1086; ORF1087; ORF1091; ORF1092; ORF1094; ORF1096; ORF1100; ORF1104; ORF1111; ORF11112; ORF1114; ORF1117; ORF1122; ORF1125 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* transmembrane polypeptide or one of its representative fragments, having at least 7 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF17; ORF52; ORF68; ORF83; ORF87; ORF109; ORF112; ORF113; ORF120; ORF121; ORF127; ORF153; ORF204; ORF211; ORF218; ORF223; ORF275; ORF277; ORF295; ORF300; ORF302; ORF306; ORF327; ORF335; ORF342; ORF343; ORF347; ORF349; ORF361; ORF363; ORF369; ORF380; ORF388; ORF389; ORF397; ORF415; ORF432; ORF439; ORF446; ORF449; ORF472; ORF478; ORF500; ORF522; ORF524; ORF567; ORF575; ORF602; ORF606; ORF609; ORF636; ORF639; ORF643; ORF653; ORF668; ORF692; ORF702; ORF704; ORF713; ORF720; ORF778; ORF784; ORF800; ORF836; ORF838; ORF842; ORF864; ORF867; ORF883; ORF901; ORF916; ORF932; ORF934; ORF940; ORF942; ORF950; ORF956; ORF971; ORF973; ORF976; ORF988; ORF994; ORF1018; ORF1028; ORF1035; ORF1037; ORF1044; ORF1055; ORF1057; ORF1068; ORF1069; ORF1070; ORF1072; ORF1082; ORF1088; ORF1105; ORF1132; ORF1135 and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* surface exposed polypeptide (e.g., an outer membrane protein) or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

ORF 15, ORF 25, ORF 26, ORF 27, ORF 28, ORF 29, ORF 30, ORF 31, ORF 32, ORF 33, ORF 35, ORF 36, ORF 1257, ORF 280, ORF 291, ORF 314, ORF 354, ORF 380, ORF 1266, ORF 466, ORF 467, ORF 468, ORF 469, ORF 470, ORF 472, ORF 474, ORF 476, ORF 477, ORF 478, ORF 479, ORF 480, ORF 482, ORF 483, ORF 485, ORF 486, ORF 500, ORF 501, ORF 503, ORF 504, ORF 505, ORF 506, ORF 507, ORF 1268, ORF 1269, ORF 543, ORF 544, ORF 578, ORF 579, ORF 580, ORF 581, ORF 595, ORF 596, ORF 597, ORF 1271, ORF 633, ORF 637, ORF 699, ORF 706, ORF 737, ORF 744, ORF 1273, ORF 751, ORF 775, ORF 776, ORF 777, ORF 793, ORF 815, ORF 830, ORF 1221, ORF 849, ORF 851, ORF 852, ORF 874, ORF 891, ORF 922, ORF 940, ORF 1231, ORF 1281, ORF 1035, ORF 1079, ORF 1087, ORF 1108, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* lipoprotein or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

ORF 3, ORF 10, ORF 11, ORF 16, ORF 1254, ORF 1255, ORF 38, ORF 1256, ORF 62, ORF 85, ORF 1258, ORF 115, ORF 1151, ORF 151, ORF 1259, ORF 173, ORF 1261, ORF 186, ORF 194, ORF 205, ORF 214, ORF 216, ORF 217, ORF 238, ORF 1177, ORF 280, ORF 291, ORF 317, ORF 327, ORF 354, ORF 364, ORF 367, ORF 414, ORF 432, ORF 1192, ORF 460, ORF 1267, ORF 1268, ORF 520, ORF 536, ORF 1270, ORF 576, ORF 597, ORF 603, ORF 609, ORF 637, ORF 1272, ORF 652, ORF 1213, ORF 699, ORF 705, ORF 706, ORF 708, ORF 711, ORF 727, ORF 1274, ORF 800, ORF 814, ORF 825, ORF 829, ORF 830, ORF 831, ORF 844, ORF 849, ORF 1275, ORF 1276, ORF 1277, ORF 872, ORF 878, ORF 880, ORF 891, ORF 892, ORF 1278, ORF 1279, ORF 1280, ORF 941, ORF 942, ORF 1282, ORF 1283, ORF 952, ORF 988, ORF 998, ORF 1009, ORF 1285, ORF

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1235, ORF 1028, ORF 1056, ORF 1070, ORF 1287, ORF 1087, ORF 1288, ORF 1289, ORF 1098, ORF 1246, ORF 1291, ORF 1108, ORF 1109, ORF 1112, ORF 1133, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide involved in lipopolysaccharide (LPS) biosynthesis, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 316, ORF 564, ORF 610, ORF 647, ORF 1211, ORF 688, ORF 924, and one of their representative fragments.

Preferably the invention relates to additional LPS-related nucleotide sequences according to the invention, characterized in that they encode:

- (a) a Chlamydia pneumoniae KDO (3-deoxy-D-manno-octulosonic acid)-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 177, ORF 1156, ORF 245, ORF 767, and one of their representative fragments;
- (b) a *Chlamydia pneumoniae* phosphomannomutase-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 74, and one of its representative fragments;
- (c) a Chlamydia pneumoniae phosphoglucomutase-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the 20 following sequences: ORF 1286, ORF 1039, and one of their representative fragments; and
  - (d) a Chlamydia pneumoniae lipid A component-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 689, ORF 690, ORF 691, ORF 1037, and one of their representative fragments.
  - Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide containing RGD (Arg-Gly-Asp) attachment sites or one of its representative fragments.
    - (a) RGD-containing proteins that are outer membrane proteins, are more likely to play a role in cell attachment. ORFs that encoded a protein containing an RGD sequence and also were classified as outer membrane proteins are ORF 468 and its representative fragments.
    - (b) An RGD-encoding ORF that showed homology to cds1, cds2, and copN type III virulence loci in *Chlamydia psittaci* (Hsia, R. et al. (1997), Type III secretion genes identity a putative virulence locus of Chlamydia. Molecular Microbiology 25:351-359) is ORF 350, and its representative fragments.

(c) The outer membrane of Chlamydia is made of cysteine-rich proteins that form a network of both intra and inter molecular disulfide links. This contributes to the integrity of the membrane since Chlamydia lacks the peptidoglycan layer that other gram-negative bacteria have. Cysteine-rich proteins that have the RGD sequence are also considered to be potential vaccine candidates. Cysteine-rich proteins were defined as proteins that had more than 3.0% cysteine in their primary amino acid sequence, above the mean genomic ORF cysteine content. The corresponding ORFs are: ORF 1290, ORF 1294, ORF 1296, and one of their representative fragments.

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(d) The outer membrane of Chlamydia may also contain small proteins that have cysteines in their N- and C-terminus that may contribute to the network formed by disulfide linkages. These proteins may be anchored in the outer membrane via their N-terminus and may have their C-terminus exposed, which then can interact with the host cells. Alternatively, these proteins may be anchored in the outer membrane via both N-and C-terminus and may have regions in the middle that may be exposed which can in turn interact with the host cells. ORFs encoding polypeptides that contain cysteines in their first 30 amino acids and also contain an RGD sequence are: ORF 105, ORF 106, ORF 114, ORF 170, ORF 171, ORF 1264, ORF 268, ORF 1265, ORF 350, ORF 393, ORF 394, ORF 451, ORF 452, ORF 453, ORF 473, ORF 499, ORF 515, ORF 519, ORF 525, ORF 526, ORF 538, ORF 611, ORF 645, ORF 686, ORF 700, ORF 746, ORF 755, ORF 756, ORF 757, ORF 789, ORF 814, ORF 855, ORF 856, ORF 878, ORF 957, ORF 958, ORF 989, ORF 1290, and one of their

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(e) RGD-containing ORFs homologous to RGD-containing ORFs from *Chlamydia* trachomatis are:

ORF 114, ORF 468, ORF 755, ORF 756, ORF 757, ORF 855, ORF 856, ORF 905, ORF 913, ORF 914, ORF 915, and one of their representative fragments.

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Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* Type III or other, non-type III secreted polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

representative fragments.

35 ORF 25, ORF 28, ORF 29, ORF 33, ORF 308, ORF 309, ORF 343, ORF 344, ORF 345, ORF 367, ORF 414, ORF 415, ORF 480, ORF 550, ORF 579, ORF 580, ORF 581, ORF 597, ORF 699, ORF 744, ORF 751, ORF 776, ORF 866, ORF 874, ORF 883, ORF 884, ORF 888, ORF 891, ORF 1293,

and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* cell wall anchored surface polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 267, ORF 271, ORF 419, ORF 590, ORF 932, ORF 1292, ORF 1295, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode Chlamydia pneumoniae polypeptides not found in Chlamydia trachomatis (Blastp. P>e<sup>-10</sup>), said nucleotide sequences comprising a nucleotide sequence chosen from 10 the following sequences: ORF 7, ORF 8, ORF 9, ORF 16, ORF 17, ORF 18, ORF 19, ORF 20, ORF 21, ORF 22, ORF 1254, ORF 23, ORF 1255, ORF 24, ORF 1139, ORF 1140, ORF 46, ORF 47, ORF 51, ORF 60, ORF 1256, ORF 61, ORF 62, ORF 63, ORF 64, ORF 1257, ORF 65, ORF 66, ORF 67, ORF 68, ORF 1143, ORF 1145, ORF 83, ORF 84, ORF 1146, ORF 85, ORF 86, ORF 87, ORF 1258. ORF 116, ORF 117, ORF 125, ORF 1148, ORF 143, ORF 1150, ORF 1151, ORF 144, ORF 145, ORF 15 147, ORF 148, ORF 149, ORF 150, ORF 152, ORF 1259, ORF 162, ORF 166, ORF 1154, ORF 167, ORF 1261, ORF 1156, ORF 1157, ORF 178, ORF 179, ORF 1158, ORF 182, ORF 183, ORF 184, ORF 185, ORF 1159, ORF 186, ORF 1160, ORF 187, ORF 188, ORF 189, ORF 190, ORF 1161, ORF 1162. ORF 191, ORF 192, ORF 194, ORF 195, ORF 1163, ORF 196, ORF 201, ORF 202, ORF 209, ORF 212, ORF 221, ORF 224, ORF 1167, ORF 226, ORF 227, ORF 228, ORF 229, ORF 230, ORF 20 231, ORF 232, ORF 1169, ORF 1170, ORF 1171, ORF 234, ORF 235, ORF 236, ORF 1172, ORF 243, ORF 251, ORF 252, ORF 1176, ORF 253, ORF 255, ORF 254, ORF 256, ORF 1177, ORF 1178, ORF 262, ORF 263, ORF 1264, ORF 278, ORF 279, ORF 1180, ORF 280, ORF 290, ORF 291, ORF 292, ORF 296, ORF 1181, ORF 297, ORF 298, ORF 300, ORF 1265, ORF 322, ORF 324, ORF 325, ORF 370, ORF 1186, ORF 371, ORF 372, ORF 1187, ORF 373, ORF 378, ORF 1266, ORF 382, ORF 25 383, ORF 384, ORF 385, ORF 386, ORF 1188, ORF 1189, ORF 391, ORF 392, ORF 398, ORF 400, ORF 403, ORF 1191, ORF 423, ORF 435, ORF 445, ORF 450, ORF 1193, ORF 456, ORF 460, ORF 461, ORF 465, ORF 1196, ORF 471, ORF 473, ORF 475, ORF 481, ORF 484, ORF 487, ORF 488, ORF 489, ORF 490, ORF 491, ORF 492, ORF 493, ORF 494, ORF 495, ORF 496, ORF 497, ORF 498, ORF 499, ORF 502, ORF 1267, ORF 1268, ORF 508, ORF 510, ORF 509, ORF 512, ORF 515, 30 ORF 519, ORF 1197, ORF 521, ORF 1198, ORF 522, ORF 524, ORF 528, ORF 534, ORF 537, ORF 1269, ORF 1270, ORF 548, ORF 551, ORF 557, ORF 1201, ORF 1203, ORF 562, ORF 566, ORF 593, ORF 595, ORF 600, ORF 1271, ORF 604, ORF 611, ORF 612, ORF 614, ORF 616, ORF 625, ORF 627, ORF 628, ORF 629, ORF 631, ORF 641, ORF 1272, ORF 648, ORF 1212, ORF 663, ORF 685, ORF 707, ORF 714, ORF 715, ORF 716, ORF 717, ORF 722, ORF 746, ORF 1273, ORF 761, 35 ORF 764, ORF 770, ORF 1217, ORF 783, ORF 1274, ORF 803, ORF 815, ORF 1220, ORF 835, ORF 1221, ORF 844, ORF 845, ORF 846, ORF 847, ORF 848, ORF 849, ORF 850, ORF 851, ORF 1275, ORF 852, ORF 862, ORF 1276, ORF 1277, ORF 873, ORF 1223, ORF 892, ORF 919, ORF 1225, ORF 1278, ORF 926, ORF 1228, ORF 1229, ORF 1230, ORF 1279, ORF 1281, ORF 1282, ORF 1283, ORF 948, ORF 950, ORF 949, ORF 951, ORF 980, ORF 982, ORF 1233, ORF 999, ORF 1000, ORF 1001, ORF 1002, ORF 1008, ORF 1285, ORF 1235, ORF 1016, ORF 1019, ORF 1027, ORF 1036, ORF 1241, ORF 1048, ORF 1049, ORF 1050, ORF 1053, ORF 1054, ORF 1064, ORF 1076, ORF 1091, ORF 1288, ORF 1093, ORF 1289, ORF 1101, ORF 1103, ORF 1245, ORF 1246, ORF 1247, ORF 1290, ORF 1291, ORF 1115, ORF 1116, ORF 1118, ORF 1120, ORF 1249, ORF 1121, ORF 1250, ORF 1126, ORF 1251, ORF 1127, ORF 1128, ORF 1130, ORF 1129, ORF 1131, ORF 1136, ORF 1253, ORF 1292, ORF 1294, ORF 1295, ORF 1296, and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, such as for example triose phosphate isomerase or pyruvate kinase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF2; ORF55; ORF56; ORF69; ORF75; ORF80; ORF100; ORF110; ORF114; ORF120; ORF121; ORF157; ORF160; ORF161; ORF172; ORF180; ORF181; ORF198; ORF200; ORF225; ORF248; ORF249; ORF276; ORF277; ORF318; ORF319; ORF320; ORF323; ORF331; ORF347; ORF375; ORF376; ORF381; ORF393; ORF394; ORF395; ORF396; ORF409; ORF446; ORF447; ORF448; ORF449; ORF513; ORF516; ORF571; ORF647; ORF662; ORF697; ORF718; ORF793; ORF794; ORF808; ORF809; ORF838; ORF839; ORF840; ORF853; ORF854; ORF918; ORF923; ORF929; ORF931; ORF938; ORF939; ORF958; ORF959; ORF960; ORF966; ORF995; ORF1021; ORF1040; ORF1041; ORF1042; ORF1085; ORF1100; ORF1102; ORF1117; ORF1118; ORF1119; ORF1120; ORF1135 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the intermediate metabolism of nucleotides or nucleic acids, such as for example CTP synthetase or GMP synthetase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF77; ORF78; ORF138; ORF189; ORF190; ORF233; ORF246; ORF338; ORF412; ORF421; 30 ORF438; ORF607; ORF648; ORF657; ORF740; ORF783; ORF967; ORF989; ORF990; ORF992; ORF1011; ORF1058; ORF1059; ORF1073; ORF1074 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of nucleic acids, such as for example 35 DNA polymerases or DNA topoisomerases, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF14; ORF59; ORF70; ORF71; ORF97; ORF113; ORF137; ORF141; ORF169; ORF285; ORF287;

ORF288; ORF313; ORF326; ORF358; ORF411; ORF443; ORF548; ORF569; ORF601; ORF651;
ORF654; ORF658; ORF659; ORF664; ORF665; ORF694; ORF698; ORF704; ORF760; ORF762;
ORF763; ORF786; ORF787; ORF788; ORF801; ORF802; ORF812; ORF819; ORF822; ORF870;
ORF897; ORF898; ORF902; ORF908; ORF916; ORF954; ORF955; ORF961; ORF983; ORF996;
ORF1007; ORF1012; ORF1013; ORF1014; ORF1015; ORF1038; ORF1137 and one of their representative fragments.

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Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of amino acids or polypeptides, such as for example serine hydroxymethyl transferase or the proteins which load amino acids onto transfer RNAs, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF99; ORF111; ORF127; ORF134; ORF140; ORF174; ORF175; ORF176; ORF353; ORF377; ORF404; ORF523; ORF539; ORF559; ORF561; ORF586; ORF598; ORF609; ORF636; ORF687; ORF700; ORF701; ORF759; ORF790; ORF857; ORF861; ORF904; ORF936; ORF952; ORF962; ORF963; ORF964; ORF965; ORF991; ORF1003; ORF1004; ORF1005; ORF1018; ORF1067; ORF1110; ORF1111; ORF1112; ORF1114; ORF1121; ORF1122; ORF1123; ORF1124; ORF1125 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of polypeptides, such as for example protein kinases or proteases, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF4; ORF44; ORF45; ORF48; ORF54; ORF112; ORF130; ORF155; ORF163; ORF212; ORF257; ORF307; ORF343; ORF405; ORF416; ORF458; ORF540; ORF541; ORF542; ORF543; ORF544; ORF560; ORF594; ORF652; ORF699; ORF723; ORF747; ORF817; ORF827; ORF871; ORF909; ORF910; ORF911; ORF912; ORF1023; ORF1051; ORF1052; ORF1081 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of fatty acids, such as for example succinyl-CoA-synthesizing proteins or phosphatidylserine synthetase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF76; ORF284; ORF308; ORF309; ORF310; ORF311; ORF312; ORF425; ORF433; ORF565; ORF688; ORF690; ORF691; ORF767; ORF797; ORF894; ORF895; ORF994; ORF1020; ORF1030; ORF1033; ORF1034; ORF1046; ORF1047; ORF1057 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its

representative fragments which is involved in the synthesis of the wall, such as for example KDO transferase, and the proteins responsible for the attachment of certain sugars onto the exposed proteins, and in that they comprise a nucleotide sequence chosen from the following sequences: ORF49; ORF50; ORF177; ORF178; ORF245; ORF610; ORF972; ORF974; ORF978; ORF1037 and

5 one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the transcription, translation and/or maturation process, such as for example initiation factors, RNA polymerases or certain chaperone proteins, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF90; ORF92; ORF131; ORF151; ORF199; ORF333; ORF334; ORF336; ORF379; ORF589; ORF590; ORF619; ORF630; ORF649; ORF739; ORF741; ORF806; ORF821; ORF843; ORF968; ORF971; ORF1061 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* ribosomal polypeptide or one of its representative fragments, such as for example the ribosomal proteins L21, L27 and S10, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF93; ORF94; ORF95; ORF136; ORF259; ORF332; ORF348; ORF583; ORF584; ORF588; ORF591; ORF592; ORF663; ORF666; ORF667; ORF669; ORF670; ORF671; ORF672; ORF673; ORF674; ORF675; ORF676; ORF677; ORF678; ORF679; ORF680; ORF681; ORF683; ORF684; ORF738; ORF781; ORF1008; ORF1024; ORF1025; ORF1066 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* transport polypeptide or one of its representative fragments, such as for example the proteins for transporting amino acids, sugars and certain oligopeptides, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF40; ORF41; ORF52; ORF105; ORF106; ORF107; ORF109; ORF133; ORF210; ORF211; ORF214; ORF215; ORF216; ORF217; ORF218; ORF219; ORF220; ORF223; ORF242; ORF260; ORF293; ORF299; ORF366; ORF369; ORF575; ORF602; ORF638; ORF639; ORF640; ORF643; ORF653; ORF702; ORF703; ORF724; ORF732; ORF855; ORF856; ORF901; ORF906; ORF933; ORF942; ORF1043; ORF1086; ORF1105 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the virulence process, such as for example the proteins analogous to the *Escherichia coli* vacB protein, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF546; ORF550; ORF778; ORF779; ORF886 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the secretory system and/or which is secreted, such as for example proteins homologous to proteins in the secretory system of certain bacteria such as the Salmonellae or the Yersiniae, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF751; ORF874; ORF875; ORF876; ORF883; ORF884; ORF885 and one of their representative fragments.

Preferably, the invention also relates to a nucleotide sequence according to the invention, characterized in that they encode a polypeptide specific to *Chlamydia pneumoniae* or one of its representative fragments (with a Blast E value of >10<sup>-5</sup>), and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF7; ORF8; ORF17; ORF18; ORF19; ORF20; ORF22; ORF23; ORF24; ORF51; ORF60; ORF63;
15 ORF65; ORF66; ORF67; ORF83; ORF84; ORF86; ORF87; ORF125; ORF143; ORF144; ORF179;
ORF182; ORF184; ORF185; ORF187; ORF221; ORF252; ORF254;; ORF278; ORF279; ORF387;
ORF388; ORF397; ORF1048; ORF1049; ORF1050; ORF1128; ORF1130; ORF1131 and one of their representative fragments.

Also forming part of the invention are polypeptides encoded by the polynucleotides of the invention, as well as fusion polypeptides comprising such polypeptides. In one embodiment, the polypeptides and fusion polypeptides immunoreact with seropositive serum of an individual infected with *Chlamydia pneumoniae*. For example, described below, are polypeptide sequences exhibiting particularly preferable characteristics. For each group of preferred polypeptides described below, it is to be understood that in addition to the individual polypeptides listed, in instances wherein such polypeptides are encoded as part of "combined" ORFs, such "combined" polypeptides are also to be included within the preferred group.

The subject of the invention is also a polypeptide according to the invention, characterized in that it is a polypeptide of the cellular envelope, preferably of the outer cellular envelope, of *Chlamydia pneumoniae* or one of its representative fragments. According to the invention, the said polypeptide is preferably chosen from the polypeptides having the following sequences:

SEQ ID No. 15; SEQ ID No. 25; SEQ ID No. 26; SEQ ID No. 27; SEQ ID No. 28; SEQ ID No. 29; SEQ ID No. 30; SEQ ID No. 31; SEQ ID No. 32; SEQ ID No. 33; SEQ ID No. 35; SEQ ID No. 68; SEQ ID No. 124; SEQ ID No. 275; SEQ ID No. 291; SEQ ID No. 294; SEQ ID No. 327; SEQ ID No. 342; SEQ ID No. 364; SEQ ID No. 374; SEQ ID No. 380; SEQ ID No. 414; SEQ ID No. 439; SEQ ID No. 466; SEQ ID No. 467; SEQ ID No. 468; SEQ ID No. 469; SEQ ID No. 470; SEQ ID No. 472; SEQ ID No. 474; SEQ ID No. 476; SEQ ID No. 477; SEQ ID No. 478; SEQ ID No. 479;

SEQ ID No. 480; SEQ ID No. 482; SEQ ID No. 485; SEQ ID No. 500; SEQ ID No. 501;
SEQ ID No. 503; SEQ ID No. 504; SEQ ID No. 505; SEQ ID No. 506; SEQ ID No. 520; SEQ ID No. 578; SEQ ID No. 580; SEQ ID No. 581; SEQ ID No. 595; SEQ ID No. 596; SEQ ID No. 597;
SEQ ID No. 737; SEQ ID No. 830; SEQ ID No. 834; SEQ ID No. 836; SEQ ID No. 893; SEQ ID No. 917; SEQ ID No. 932; SEQ ID No. 976; SEQ ID No. 1035; SEQ ID No. 1045; SEQ ID No. 1090 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a Chlamydia pneumoniae transmembrane polypeptide or one of its representative fragments, having between 1 and 3 transmembrane domains, and in that it is chosen 10 from the polypeptides having the following sequences: SEQ ID No. 2; SEQ ID No. 3; SEQ ID No. 6; SEQ ID No. 9; SEQ-ID No. 10; SEQ-ID No. 11; SEQ ID No. 13; SEQ ID No. 14; SEQ ID No. 16; SEQ ID No. 18; SEQ ID No. 19; SEQ ID No. 20; SEO ID - No. 21; SEO ID No. 22; SEQ ID No. 25; SEQ ID No. 27; SEQ ID No. 28; SEQ ID No. 29; SEQ ID No. 30; SEQ ID No. 31; SEQ ID No. 32; SEQ ID No. 33; SEQ ID No. 34; 15 SEO ID No. 35; SEQ ID No. 37; SEQ ID No. 39; SEQ ID No. 41; SEQ ID No. 42; SEQ ID No. 44; SEQ ID No. 45; SEQ ID No. 46; SEQ ID No. 47; SEQ ID No. 48; SEQ ID No. 49; SEQ ID No. 50; SEQ ID No. 53; SEQ ID No. 54; SEQ ID No. 56; SEQ ID No. 57; SEQ ID No. 59; SEQ ID No. 60; SEQ ID No. 61; SEQ ID No. 62; SEQ ID No. 63; SEQ ID No. 64; SEQ ID No. 65; SEQ ID No. 66; SEQ ID No. 69;; SEQ ID No. 72; SEQ ID No. 73; SEQ ID 20 No. 74; SEQ ID No. 76; SEQ ID No. 77; SEQ ID No. 78; SEQ ID No. 79; SEQ ID No. 80; SEQ ID No. 82; SEQ ID No. 84; SEQ ID No. 85; SEQ ID No. 86; SEQ ID No. 88; SEQ ID No. 89; SEQ ID No. 90; SEQ ID No. 91; SEQ ID No. 92; SEQ ID No. 93; SEQ ID No. 95; SEQ ID No. 96; SEQ ID No. 98; SEQ ID No. 99; SEQ ID No. 100; SEQ ID No. 101; SEQ ID No. 102; SEQ ID No. 103; SEQ ID No. 104; SEQ ID No. 105; SEQ ID No. 106; SEQ ID No. 107; 25 SEQ ID No. 108; SEQ ID No. 114; SEQ ID No. 117; SEQ ID No. 118; SEQ ID No. 122; SEQ ID No. 123; SEQ ID No. 124; SEQ ID No. 125; SEQ ID No. 129; SEQ ID No. 130; SEQ ID No. 131; SEQ ID No. 132; SEQ ID No. 133; SEQ ID No. 134; SEQ ID No. 135; SEQ ID No. 137; SEQ ID No. 138; SEQ ID No. 139; SEQ ID No. 140; SEQ ID No. 141; SEQ ID No. 142; SEQ ID No. 143; SEQ ID No. 145; SEQ ID No. 146; SEQ ID No. 147; SEQ ID No. 150; SEQ ID No. 151; SEQ ID 30 No. 152; SEQ ID No. 156; SEQ ID No. 157; SEQ ID No. 158; SEQ ID No. 159; SEQ ID No. 160; SEQ ID No. 161; SEQ ID No. 162; SEQ ID No. 164; SEQ ID No. 166; SEQ ID No. 167; SEQ ID No. 170; SEQ ID No. 173; SEQ ID No. 175; SEQ ID No. 176; SEQ ID No. 178; SEQ ID No. 179; SEQ ID No. 180; SEQ ID No. 182; SEQ ID No. 183; SEQ ID No. 184; SEQ ID No. 185; SEQ ID No. 186; SEQ ID No. 187; SEQ ID No. 188; SEQ ID No. 189; SEQ ID No. 190; SEQ ID No. 191; 35 SEQ ID No. 192; SEQ ID No. 194; SEQ ID No. 195; SEQ ID No. 196; SEQ ID No. 197; SEQ ID No. 198; SEQ ID No. 199; SEQ ID No. 200; SEQ ID No. 201; SEQ ID No. 202; SEQ ID No. 205; SEQ ID No. 207; SEQ ID No. 208; SEQ ID No. 209; SEQ ID No. 210; SEQ ID No. 212; SEQ ID

No. 215; SEQ ID No. 219; SEQ ID No. 220; SEQ ID No. 224; SEQ ID No. 226; SEQ ID No. 227; SEQ ID No. 228; SEQ ID No. 231; SEQ ID No. 232; SEQ ID No. 233; SEQ ID No. 234; SEO ID No. 235; SEQ ID No. 236; SEQ ID No. 238; SEQ ID No. 239; SEQ ID No. 240; SEQ ID No. 241; SEQ ID No. 242; SEQ ID No. 244; SEQ ID No. 247; SEQ ID No. 251; SEQ ID No. 252; 5 SEQ ID No. 253; SEQ ID No. 255; SEQ ID No. 256; SEQ ID No. 257; SEQ ID No. 258; SEQ ID No. 260; SEQ ID No. 262; SEQ ID No. 263; SEQ ID No. 266; SEQ ID No. 267; SEQ ID No. 268; SEQ ID No. 269; SEQ ID No. 270; SEQ ID No. 273; SEQ ID No. 274; SEQ ID No. 276; SEQ ID No. 278; SEQ ID No. 279; SEQ ID No. 280; SEQ ID No. 281; SEQ ID No. 282; SEQ ID No. 283; SEQ ID No. 284; SEQ ID No. 286; SEQ ID No. 287; SEQ ID No. 289; SEQ ID No. 290; SEQ ID 10 No. 291; SEQ ID No. 293; SEQ ID No. 294; SEQ ID No. 297; SEQ ID No. 304; SEQ ID No. 305; SEO ID No. 307; SEQ ID No. 308; SEQ ID No. 309; SEQ ID No. 310; SEQ ID No. 311; SEQ ID No. 313; SEQ ID No. 314; SEQ ID No. 315; SEQ ID No. 316; SEQ ID No. 318; SEQ ID No. 319; SEQ ID No. 320; SEQ ID No. 321; SEQ ID No. 322; SEQ ID No. 323; SEQ ID No. 324; SEQ ID No. 325; SEQ ID No. 326; SEQ ID No. 331; SEQ ID No. 332; SEQ ID No. 336; SEQ ID No. 338; 15 SEQ ID No. 339; SEQ ID No. 341; SEQ ID No. 344; SEQ ID No. 345; SEQ ID No. 346; SEQ ID No. 350; SEQ ID No. 352; SEQ ID No. 353; SEQ ID No. 356; SEQ ID No. 357; SEQ ID No. 358; SEQ ID No. 359; SEQ ID No. 360; SEQ ID No. 362; SEQ ID No. 365; SEQ ID No. 366; SEQ ID No. 367; SEO ID No. 370; SEO ID No. 372; SEQ ID No. 373; SEQ ID No. 376; SEQ ID No. 377; SEQ ID No. 378; SEQ ID No. 379; SEQ ID No. 381; SEQ ID No. 382; SEQ ID No. 383; SEQ ID 20 No. 384; SEQ ID No. 385; SEQ ID No. 386; SEQ ID No. 387; SEQ ID No. 390; SEQ ID No. 392; SEQ ID No. 393; SEQ ID No. 394; SEQ ID No. 396; SEQ ID No. 398; SEQ ID No. 399; SEQ ID No. 400; SEQ ID No. 404; SEQ ID No. 408; SEQ ID No. 410; SEQ ID No. 411; SEQ ID No. 413; SEQ ID No. 416; SEQ ID No. 417; SEQ ID No. 418; SEQ ID No. 420; SEQ ID No. 422; SEQ ID No. 424; SEQ ID No. 427; SEQ ID No. 428; SEQ ID No. 429; SEQ ID No. 430; SEQ ID No. 431; 25 SEO ID No. 433; SEO ID No. 434; SEQ ID No. 437; SEQ ID No. 440; SEQ ID No. 441; SEQ ID No. 442; SEO ID No. 443; SEQ ID No. 444; SEQ ID No. 445; SEQ ID No. 447; SEQ ID No. 450; SEQ ID No. 451; SEQ ID No. 452; SEQ ID No. 455; SEQ ID No. 456; SEQ ID No. 459; SEQ ID No. 460; SEQ ID No. 461; SEQ ID No. 462; SEQ ID No. 463; SEQ ID No. 464; SEQ ID No. 465; SEQ ID No. 467; SEQ ID No. 469; SEQ ID No. 471; SEQ ID No. 474; SEQ ID No. 475; SEQ ID 30 No. 476; SEQ ID No. 477; SEQ ID No. 479; SEQ ID No. 482; SEQ ID No. 483; SEQ ID No. 484; SEQ ID No. 485; SEQ ID No. 486; SEQ ID No. 487; SEQ ID No. 488; SEQ ID No. 491; SEQ ID No. 493; SEQ ID No. 494; SEQ ID No. 497; SEQ ID No. 498; SEQ ID No. 499; SEQ ID No. 503; SEQ ID No. 508; SEQ ID No. 509; SEQ ID No. 510; SEQ ID No. 512; SEQ ID No. 514; SEQ ID No. 515; SEQ ID No. 516; SEQ ID No. 517; SEQ ID No. 518; SEQ ID No. 520; SEQ ID No. 521; 35 SEO ID No. 523; SEO ID No. 525; SEQ ID No. 527; SEQ ID No. 528; SEQ ID No. 529; SEQ ID No. 530; SEQ ID No. 531; SEQ ID No. 533; SEQ ID No. 534; SEQ ID No. 535; SEQ ID No. 536; SEQ ID No. 537; SEQ ID No. 540; SEQ ID No. 541; SEQ ID No. 543; SEQ ID No. 544; SEQ ID

No. 545; SEO ID No. 546; SEO ID No. 548; SEO ID No. 549; SEO ID No. 551; SEO ID No. 553; SEQ ID No. 554; SEQ ID No. 555; SEQ ID No. 556; SEQ ID No. 557; SEQ ID No. 558; SEO ID No. 559; SEQ ID No. 560; SEQ ID No. 562; SEQ ID No. 563; SEQ ID No. 564; SEQ ID No. 565; SEQ ID No. 566; SEQ ID No. 569; SEQ ID No. 571; SEQ ID No. 573; SEQ ID No. 576; 5 SEO ID No. 577; SEQ ID No. 581; SEQ ID No. 583; SEQ ID No. 584; SEQ ID No. 585; SEQ ID No. 586; SEQ ID No. 588; SEQ ID No. 591; SEQ ID No. 592; SEQ ID No. 594; SEQ ID No. 595; SEQ ID No. 596; SEQ ID No. 597; SEQ ID No. 599; SEQ ID No. 600; SEQ ID No. 603; SEQ ID No. 605; SEQ ID No. 608; SEQ ID No. 614; SEQ ID No. 615; SEQ ID No. 620; SEQ ID No. 621; SEQ ID No. 622; SEQ ID No. 623; SEQ ID No. 624; SEQ ID No. 625; SEQ ID No. 629; SEQ ID 10 No. 630; SEQ ID No. 631; SEQ ID No. 633; SEQ ID No. 634; SEQ ID No. 637; SEQ ID No. 642; SEQ ID No. 644; SEQ ID No. 645; SEQ-ID No. 647; SEQ ID No. 648; SEQ ID No. 652; SEQ ID No. 654; SEQ ID No. 655; SEQ ID No. 657; SEQ ID No. 658; SEQ ID No. 659; SEQ ID No. 660; SEQ ID No. 661; SEQ ID No. 664; SEQ ID No. 665; SEQ ID No. 666; SEQ ID No. 667; SEQ ID No. 670; SEQ ID No. 671; SEQ ID No. 672; SEQ ID No. 673; SEQ ID No. 674; SEQ ID No. 676; 15 SEQ ID No. 679; SEQ ID No. 681; SEQ ID No. 684; SEQ ID No. 687; SEQ ID No. 688; SEQ ID No. 689; SEO ID No. 690; SEO ID No. 693; SEO ID No. 694; SEQ ID No. 695; SEQ ID No. 696; SEQ ID No. 697; SEQ ID No. 698; SEQ ID No. 699; SEQ ID No. 700; SEQ ID No. 701; SEQ ID No. 703; SEQ ID No. 705; SEQ ID No. 706; SEQ ID No. 707; SEQ ID No. 708; SEQ ID No. 710; SEQ ID No. 712; SEQ ID No. 715; SEQ ID No. 716; SEQ ID No. 717; SEQ ID No. 718; SEQ ID 20 No. 719; SEQ ID No. 721; SEQ ID No. 722; SEQ ID No. 723; SEQ ID No. 725; SEQ ID No. 726; SEQ ID No. 727; SEQ ID No. 728; SEQ ID No. 729; SEQ ID No. 730; SEQ ID No. 731; SEQ ID No. 733; SEQ ID No. 736; SEQ ID No. 737; SEQ ID No. 738; SEQ ID No. 740; SEQ ID No. 741; SEQ ID No. 742; SEQ ID No. 743; SEQ ID No. 747; SEQ ID No. 748; SEQ ID No. 750; SEQ ID No. 752; SEQ ID No. 754; SEQ ID No. 755; SEQ ID No. 756; SEQ ID No. 757; SEQ ID No. 759; 25 SEO ID No. 760; SEO ID No. 761; SEQ ID No. 762; SEQ ID No. 763; SEQ ID No. 764; SEQ ID No. 765; SEQ ID No. 766; SEQ ID No. 767; SEQ ID No. 768; SEQ ID No. 772; SEQ ID No. 774; SEQ ID No. 775; SEQ ID No. 777; SEQ ID No. 781; SEQ ID No. 783; SEQ ID No. 788; SEQ ID No. 791; SEQ ID No. 792; SEQ ID No. 793; SEQ ID No. 794; SEQ ID No. 795; SEQ ID No. 796; SEQ ID No. 797; SEQ ID No. 798; SEQ ID No. 799; SEQ ID No. 802; SEQ ID No. 803; SEQ ID 30 No. 806; SEQ ID No. 807; SEQ ID No. 808; SEQ ID No. 809; SEQ ID No. 810; SEQ ID No. 811; SEO ID No. 813; SEO ID No. 814; SEO ID No. 815; SEO ID No. 816; SEQ ID No. 817; SEQ ID No. 819; SEQ ID No. 820; SEQ ID No. 821; SEQ ID No. 823; SEQ ID No. 824; SEQ ID No. 827; SEQ ID No. 829; SEQ ID No. 830; SEQ ID No. 831; SEQ ID No. 833; SEQ ID No. 834; SEQ ID No. 835; SEQ ID No. 837; SEQ ID No. 844; SEQ ID No. 845; SEQ ID No. 846; SEQ ID No. 847; 35 SEQ ID No. 848; SEQ ID No. 849; SEQ ID No. 850; SEQ ID No. 851; SEQ ID No. 852; SEQ ID No. 854; SEQ ID No. 855; SEQ ID No. 856; SEQ ID No. 857; SEQ ID No. 859; SEQ ID No. 860; SEQ ID No. 862; SEQ ID No. 865; SEQ ID No. 866; SEQ ID No. 868; SEQ ID No. 869; SEQ ID

No. 870; SEQ ID No. 871; SEQ ID No. 872; SEQ ID No. 874; SEQ ID No. 877; SEQ ID No. 878; SEO ID No. 879; SEO ID No. 880; SEQ ID No. 881; SEQ ID No. 882; SEQ ID No. 884; SEO ID No. 885; SEQ ID No. 888; SEQ ID No. 889; SEQ ID No. 890; SEQ ID No. 891; SEQ ID No. 892; SEQ ID No. 894; SEQ ID No. 895; SEQ ID No. 896; SEQ ID No. 897; SEQ ID No. 899; 5 SEQ ID No. 900; SEQ ID No. 902; SEQ ID No. 903; SEQ ID No. 904; SEQ ID No. 905; SEQ ID No. 909; SEQ ID No. 910; SEQ ID No. 912; SEQ ID No. 913; SEQ ID No. 914; SEQ ID No. 915; SEO ID No. 917; SEQ ID No. 918; SEQ ID No. 919; SEQ ID No. 921; SEQ ID No. 923; SEQ ID No. 924; SEQ ID No. 926; SEQ ID No. 927; SEQ ID No. 928; SEQ ID No. 929; SEQ ID No. 930; SEQ ID No. 931; SEQ ID No. 937; SEQ ID No. 938; SEQ ID No. 939; SEQ ID No. 941; SEO ID 10 No. 943; SEQ ID No. 948; SEQ ID No. 951; SEQ ID No. 952; SEQ ID No. 953; SEQ ID No. 958; SEO ID No. 960; SEO ID No. 963; SEQ ID No. 964; SEQ ID No. 965; SEQ ID No. 968; SEQ ID No. 970; SEQ ID No. 974; SEQ ID No. 975; SEQ ID No. 977; SEQ ID No. 979; SEQ ID No. 980; SEO ID No. 981; SEO ID No. 983; SEQ ID No. 984; SEQ ID No. 985; SEQ ID No. 987; SEQ ID No. 989; SEQ ID No. 992; SEQ ID No. 993; SEQ ID No. 997; SEQ ID No. 998; SEQ ID No. 999; 15 SEQ ID No. 1001; SEQ ID No. 1002; SEQ ID No. 1004; SEQ ID No. 1005; SEQ ID No. 1009; SEO ID No. 1013; SEQ ID No. 1014; SEQ ID No. 1015; SEQ ID No. 1016; SEQ ID No. 1019; SEQ ID No. 1021; SEQ ID No. 1023; SEQ ID No. 1024; SEQ ID No. 1029; SEQ ID No. 1031; SEQ ID No. 1033; SEQ ID No. 1034; SEQ ID No. 1039; SEQ ID No. 1041; SEQ ID No. 1042; SEO ID No. 1045; SEO ID No. 1047; SEQ ID No. 1049; SEQ ID No. 1051; SEQ ID No. 1052; 20 SEQ ID No. 1053; SEQ ID No. 1054; SEQ ID No. 1056; SEQ ID No. 1059; SEQ ID No. 1061; SEQ ID No. 1062; SEQ ID No. 1063; SEQ ID No. 1064; SEQ ID No. 1065; SEQ ID No. 1067; SEQ ID No. 1075; SEQ ID No. 1077; SEQ ID No. 1078; SEQ ID No. 1079; SEQ ID No. 1080; SEQ ID No. 1081; SEQ ID No. 1089; SEQ ID No. 1095; SEQ ID No. 1097; SEQ ID No. 1098; SEO ID No. 1099; SEO ID No. 1101; SEO ID No. 1102; SEQ ID No. 1103; SEQ ID No. 1106; 25 SEO ID No. 1107; SEO ID No. 1108; SEQ ID No. 1109; SEQ ID No. 1110; SEQ ID No. 1113; SEO ID No. 1116; SEO ID No. 1118; SEO ID No. 1119; SEQ ID No. 1121; SEQ ID No. 1123; SEO ID No. 1124; SEO ID No. 1126; SEQ ID No. 1128; SEQ ID No. 1130; SEQ ID No. 1131; SEQ ID No. 1133; SEQ ID No. 1134; SEQ ID No. 1136; SEQ ID No. 1137 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* transmembrane polypeptide or one of its respective fragments, having between 4 and 6 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 5; SEQ ID No. 7; SEQ ID No. 8; SEQ ID No. 15; SEQ ID No. 36; SEQ ID No. 38; SEQ ID No. 51; SEQ ID No. 55; SEQ ID No. 58; SEQ ID No. 67; SEQ ID No. 70; SEQ ID No. 81; SEQ ID No. 97; SEQ ID No. 110; SEQ ID No. 111; SEQ ID No. 115; SEQ ID No. 119; SEQ ID No. 126; SEQ ID No. 128; SEQ ID No. 148; SEQ ID No. 155; SEQ ID No. 163; SEQ ID

No. 165; SEQ ID No. 168; SEQ ID No. 169; SEQ ID No. 171; SEQ ID No. 172; SEQ ID No. 174; SEQ ID No. 177; SEQ ID No. 181; SEQ ID No. 193; SEQ ID No. 203; SEQ ID No. 213; SEO ID No. 214; SEQ ID No. 216; SEQ ID No. 217; SEQ ID No. 221; SEQ ID No. 222; SEQ ID No. 225; SEQ ID No. 229; SEQ ID No. 243; SEQ ID No. 246; SEQ ID No. 248; SEQ ID No. 254; 5 SEQ ID No. 261; SEQ ID No. 285; SEQ ID No. 288; SEQ ID No. 292; SEQ ID No. 296; SEQ ID No. 298; SEQ ID No. 299; SEQ ID No. 301; SEQ ID No. 303; SEQ ID No. 317; SEQ ID No. 328; SEQ ID No. 329; SEQ ID No. 351; SEQ ID No. 354; SEQ ID No. 355; SEQ ID No. 364; SEQ ID No. 371; SEO ID No. 374; SEO ID No. 375; SEO ID No. 391; SEO ID No. 395; SEO ID No. 401; SEO ID No. 403; SEQ ID No. 405; SEQ ID No. 409; SEQ ID No. 414; SEQ ID No. 419; SEQ ID 10 No. 421; SEQ ID No. 423; SEQ ID No. 425; SEQ ID No. 438; SEQ ID No. 448; SEQ ID No. 453; SEO ID No. 458; SEQ ID No. 466; SEQ ID No. 468; SEQ ID No. 470; SEQ ID No. 480; SEQ ID No. 489; SEQ ID No. 490; SEQ ID No. 496; SEQ ID No. 501; SEQ ID No. 504; SEQ ID No. 505; SEQ ID No. 506; SEQ ID No. 511; SEQ ID No. 513; SEQ ID No. 519; SEQ ID No. 526; SEQ ID No. 532; SEQ ID No. 538; SEQ ID No. 539; SEQ ID No. 547; SEQ ID No. 550; SEQ ID No. 561; 15 SEO ID No. 568; SEO ID No. 570; SEQ ID No. 574; SEQ ID No. 578; SEQ ID No. 579; SEQ ID No. 580; SEQ ID No. 582; SEQ ID No. 589; SEQ ID No. 593; SEQ ID No. 598; SEQ ID No. 601; SEO ID No. 604; SEQ ID No. 610; SEQ ID No. 613; SEQ ID No. 617; SEQ ID No. 626; SEQ ID No. 632; SEQ ID No. 635; SEQ ID No. 638; SEQ ID No. 640; SEQ ID No. 641; SEQ ID No. 646; SEQ ID No. 649; SEQ ID No. 650; SEQ ID No. 651; SEQ ID No. 686; SEQ ID No. 711; SEQ ID 20 No. 724; SEQ ID No. 732; SEQ ID No. 734; SEQ ID No. 744; SEQ ID No. 745; SEQ ID No. 749; SEQ ID No. 751; SEQ ID No. 769; SEQ ID No. 770; SEQ ID No. 771; SEQ ID No. 773; SEQ ID No. 776; SEQ ID No. 779; SEQ ID No. 780; SEQ ID No. 785; SEQ ID No. 787; SEQ ID No. 789; SEQ ID No. 801; SEQ ID No. 805; SEQ ID No. 812; SEQ ID No. 822; SEQ ID No. 825; SEQ ID No. 826; SEO ID No. 839; SEO ID No. 841; SEO ID No. 843; SEQ ID No. 853; SEQ ID No. 861; 25 SEQ ID No. 875; SEQ ID No. 876; SEQ ID No. 886; SEQ ID No. 893; SEQ ID No. 898; SEQ ID No. 906; SEQ ID No. 907; SEQ ID No. 908; SEQ ID No. 920; SEQ ID No. 922; SEQ ID No. 925; SEQ ID No. 933; SEQ ID No. 935; SEQ ID No. 936; SEQ ID No. 944; SEQ ID No. 946; SEQ ID No. 947; SEQ ID No. 954; SEQ ID No. 959; SEQ ID No. 961; SEQ ID No. 966; SEQ ID No. 967; SEQ ID No. 972; SEQ ID No. 978; SEQ ID No. 995; SEQ ID No. 996; SEQ ID No. 1000; SEQ ID 30 No. 1003; SEQ ID No. 1010; SEQ ID No. 1011; SEQ ID No. 1012; SEQ ID No. 1017; SEQ ID No. 1020; SEQ ID No. 1030; SEQ ID No. 1036; SEQ ID No. 1038; SEQ ID No. 1043; SEQ ID No. 1046; SEQ ID No. 1048; SEQ ID No. 1050; SEQ ID No. 1058; SEQ ID No. 1071; SEQ ID No. 1073; SEQ ID No. 1084; SEQ ID No. 1085; SEQ ID No. 1086; SEQ ID No. 1087; SEQ ID No. 1091; SEQ ID No. 1092; SEQ ID No. 1094; SEQ ID No. 1096; SEQ ID No. 1100; SEQ ID 35 No. 1104; SEQ ID No. 1111; SEQ ID No. 1112; SEQ ID No. 1114; SEQ ID No. 1117; SEQ ID No. 1122; SEQ ID No. 1125 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention,

characterized in that it is a *Chlamydia pneumoniae* transmembrane polypeptide or one of its representative fragments, having at least 7 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences:

SEO ID No. 17; SEQ ID No. 52; SEQ ID No. 68; SEQ ID No. 83; SEQ ID No. 87; SEQ ID No. 109; 5 SEO ID No. 112; SEQ ID No. 113; SEQ ID No. 120; SEQ ID No. 121; SEQ ID No. 127; SEQ ID No. 153; SEQ ID No. 204; SEQ ID No. 211; SEQ ID No. 218; SEQ ID No. 223; SEQ ID No. 275; SEQ ID No. 277; SEQ ID No. 295; SEQ ID No. 300; SEQ ID No. 302; SEQ ID No. 306; SEQ ID No. 327; SEO ID No. 335; SEO ID No. 342; SEQ ID No. 343; SEQ ID No. 347; SEQ ID No. 349; SEO ID No. 361; SEO ID No. 363; SEO ID No. 369; SEO ID No. 380; SEO ID No. 388; SEO ID 10 No. 389; SEO ID No. 397; SEO ID No. 415; SEO ID No. 432; SEQ ID No. 439; SEQ ID No. 446; SEQ ID No. 449; SEQ ID No. 472; SEQ ID No. 478; SEQ ID No. 500; SEQ ID No. 522; SEQ ID No. 524; SEQ ID No. 567; SEQ ID No. 575; SEQ ID No. 602; SEQ ID No. 606; SEQ ID No. 609; SEQ ID No. 636; SEQ ID No. 639; SEQ ID No. 643; SEQ ID No. 653; SEQ ID No. 668; SEQ ID No. 692; SEQ ID No. 702; SEQ ID No. 704; SEQ ID No. 713; SEQ ID No. 720; SEQ ID No. 778; 15 SEQ ID No. 784; SEQ ID No. 800; SEQ ID No. 836; SEQ ID No. 838; SEQ ID No. 842; SEQ ID No. 864; SEQ ID No. 867; SEQ ID No. 883; SEQ ID No. 901; SEQ ID No. 916; SEQ ID No. 932; SEQ ID No. 934; SEQ ID No. 940; SEQ ID No. 942; SEQ ID No. 950; SEQ ID No. 956; SEQ ID No. 971; SEO ID No. 973; SEO ID No. 976; SEQ ID No. 988; SEQ ID No. 994; SEQ ID No. 1018; SEQ ID No. 1028; SEQ ID No. 1035; SEQ ID No. 1037; SEQ ID No. 1044; SEQ ID No. 1055; 20 SEQ ID No. 1057; SEQ ID No. 1068; SEQ ID No. 1069; SEQ ID No. 1070; SEQ ID No. 1072; SEQ ID No. 1082; SEQ ID No. 1088; SEQ ID No. 1105; SEQ ID No. 1132; SEQ ID No. 1135 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia pneumoniae surface exposed polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 15, SEQ ID No. 25, SEQ ID No. 26, SEQ ID No. 27, SEQ ID No. 28, SEQ ID No. 29, SEQ ID No. 30, SEQ ID No. 31, SEQ ID No. 32, SEQ ID No. 33, SEQ ID No. 35, SEQ ID No. 36, SEQ ID No. 1257, SEQ ID No. 280, SEQ ID No. 291, SEQ ID No. 314, SEQ ID No. 354, SEQ ID No. 380, SEQ ID No. 1266, SEQ ID No. 466, SEQ ID No. 467, SEQ ID No. 468, SEQ ID No. 469, SEQ ID No. 470, SEQ ID No. 472, SEQ ID No. 474, SEQ ID No. 476, SEQ ID No. 477, SEQ ID No. 478, SEQ ID No. 479, SEQ ID No. 480, SEQ ID No. 482, SEQ ID No. 483, SEQ ID No. 485, SEQ ID No. 486, SEQ ID No. 500, SEQ ID No. 501, SEQ ID No. 503, SEQ ID No. 504, SEQ ID No. 505, SEQ ID No. 506, SEQ ID No. 507, SEQ ID No. 1268, SEQ ID No. 1269, SEQ ID No. 543, SEQ ID No. 544, SEQ ID No. 578, SEQ ID No. 579, SEQ ID No. 580, SEQ ID No. 581, SEQ ID No. 595, SEQ ID No. 596, SEQ ID No. 597, SEQ ID No. 1271, SEQ ID No. 633, SEQ ID No. 637, SEQ ID No. 699, SEQ ID No. 706, SEQ ID No. 737, SEQ ID No. 744, SEQ ID No. 1273, SEQ ID No. 751,

SEQ ID No. 775, SEQ ID No. 776, SEQ ID No. 777, SEQ ID No. 793, SEQ ID No. 815, SEQ ID No.

830, SEQ ID No. 1221, SEQ ID No. 849, SEQ ID No. 851, SEQ ID No. 852, SEQ ID No. 874, SEQ ID No. 891, SEQ ID No. 922, SEQ ID No. 940, SEQ ID No. 1231, SEQ ID No. 1281, SEQ ID No. 1035, SEQ ID No. 1079, SEQ ID No. 1087, SEQ ID No. 1108, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a Chlamydia pneumoniae lipoprotein or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEO ID No. 3, SEQ ID No. 10, SEQ ID No. 11, SEQ ID No. 16, SEQ ID No. 1254, SEQ ID No. 1255, SEQ ID No. 38, SEQ ID No. 1256, SEQ ID No. 62, SEQ ID No. 85, SEQ ID No. 1258, SEQ ID 10 No. 115, SEQ ID No. 1151, SEQ ID No. 151, SEQ ID No. 1259, SEQ ID No. 173, SEQ ID No. 1261, SEO ID No. 186, SEO ID No. 194, SEO ID No. 205, SEQ ID No. 214, SEQ ID No. 216, SEQ ID No. 217, SEQ ID No. 238, SEQ ID No. 1177, SEQ ID No. 280, SEQ ID No. 291, SEQ ID No. 317, SEQ ID No. 327, SEQ ID No. 354, SEQ ID No. 364, SEQ ID No. 367, SEQ ID No. 414, SEQ ID No. 432, SEQ ID No. 1192, SEQ ID No. 460, SEQ ID No. 1267, SEQ ID No. 1268, SEQ ID No. 520, SEQ ID 15 No. 536, SEQ ID No. 1270, SEQ ID No. 576, SEQ ID No. 597, SEQ ID No. 603, SEQ ID No. 609, SEQ ID No. 637, SEQ ID No. 1272, SEQ ID No. 652, SEQ ID No. 1213, SEQ ID No. 699, SEQ ID No. 705, SEQ ID No. 706, SEQ ID No. 708, SEQ ID No. 711, SEQ ID No. 727, SEQ ID No. 1274, SEQ ID No. 800, SEQ ID No. 814, SEQ ID No. 825, SEQ ID No. 829, SEQ ID No. 830, SEQ ID No. 831, SEQ ID No. 844, SEQ ID No. 849, SEQ ID No. 1275, SEQ ID No. 1276, SEQ ID No. 1277, SEQ 20 ID No. 872, SEQ ID No. 878, SEQ ID No. 880, SEQ ID No. 891, SEQ ID No. 892, SEQ ID No. 1278, SEQ ID No. 1279, SEQ ID No. 1280, SEQ ID No. 941, SEQ ID No. 942, SEQ ID No. 1282, SEQ ID No. 1283, SEQ ID No. 952, SEQ ID No. 988, SEQ ID No. 998, SEQ ID No. 1009, SEQ ID No. 1285, SEQ ID No. 1235, SEQ ID No. 1028, SEQ ID No. 1056, SEQ ID No. 1070, SEQ ID No. 1287, SEQ ID No. 1087, SEQ ID No. 1288, SEQ ID No. 1289, SEQ ID No. 1098, SEQ ID No. 1246, SEQ ID No. 25 1291, SEQ ID No. 1108, SEQ ID No. 1109, SEQ ID No. 1112, SEQ ID No. 1133, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia pneumoniae polypeptide involved in lipopolysaccharide (LPS) biosynthesis, and in that it is chosen from the polypeptides having the following sequences:

30 SEQ ID No. 316, SEQ ID No. 564, SEQ ID No. 610, SEQ ID No. 647, SEQ ID No. 1211, SEQ ID No. 688, SEQ ID No. 924, and one of their representative fragments.

Preferably, the invention relates to additional LPS-related polypeptides according to the invention, in that it is:

(a) a Chlamydia pneumoniae KDO (3-deoxy-D-manno-octylosonic acid)-related 35 polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 177, SEQ ID No. 1156, SEQ ID No. 245, SEQ ID No. 767, and one of their representative fragments;

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representative fragments.

- (b) a Chlamydia pneumoniae phosphomannomutase-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 74, and its representative fragment;
- (c) a Chlamydia pneumoniae phosphoglucomutase-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 1286, SEQ ID No. 1039, and its representative fragment; and
- (d) a Chlamydia pneumoniae lipid A component-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 689, SEQ ID No. 690, SEQ ID No. 691, SEQ ID No. 1037, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia pneumoniae polypeptide or one of its representative fragments that contains an RGD sequence and is also an outer membrane protein, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 468 and its representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments that contains an RGD sequence that shows homology to cds1, cds2, and copN type III virulence loci in *Chlamydia Psitacci*, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 350 and its representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia pneumoniae polypeptide or one of its representative fragments that is cysteine-rich and contains RGD sequence, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 1290, SEQ ID No. 6846, SEQ ID No. 6848, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia pneumoniae* outer membrane polypeptide that contains cysteines in their first 30 amino acids and also contain an RGD sequence, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 105, SEQ ID No. 106, SEQ ID No. 114, SEQ ID No. 170, SEQ ID No. 171, SEQ ID No. 1264, SEQ ID No. 268, SEQ ID No. 1265, SEQ ID No. 350, SEQ ID No. 393, SEQ ID No. 394, SEQ ID No. 451, SEQ ID No. 452, SEQ ID No. 453, SEQ ID No. 473, SEQ ID No. 499, SEQ ID No. 515, SEQ ID No. 519, SEQ ID No. 525, SEQ ID No. 526, SEQ ID No. 538, SEQ ID No. 611, SEQ ID No. 645, SEQ ID No. 686, SEQ ID No. 700, SEQ ID No. 746, SEQ ID No. 755, SEQ ID No. 756, SEQ ID No. 757, SEQ ID No. 789, SEQ ID No. 814, SEQ ID No. 855, SEQ ID No. 856, SEQ ID No. 878, SEQ ID No. 957, SEQ ID No. 958, SEQ ID No. 989, SEQ ID No. 1290, and one of their

Preferably, the invention relates to a polypeptide according to the invention, in that it is a

Chlamydia pneumoniae polypeptide or one of its representative fragments that contains RGD sequences homologous to Chlamydia trachomatis polypeptides containing RGD sequences, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 114, SEQ ID No. 468, SEQ ID No. 755, SEQ ID No. 756, SEQ ID No. 757, SEQ ID No.
855, SEQ ID No. 856, SEQ ID No. 905, SEQ ID No. 913, SEQ ID No. 914, SEQ ID No. 915, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia pneumoniae Type III and non-Type III secreted polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 25, SEQ ID No. 28, SEQ ID No. 29, SEQ ID No. 33, SEQ ID No. 308, SEQ ID No. 309, SEQ ID No. 343, SEQ ID No. 344, SEQ ID No. 345, SEQ ID No. 367, SEQ ID No. 414, SEQ ID No. 415, SEQ ID No. 480, SEQ ID No. 550, SEQ ID No. 579, SEQ ID No. 580, SEQ ID No. 581, SEQ ID No. 597, SEQ ID No. 699, SEQ ID No. 744, SEQ ID No. 751, SEQ ID No. 776, SEQ ID No. 866, SEQ ID No. 874, SEQ ID No. 883, SEQ ID No. 884, SEQ ID No. 888, SEQ ID No. 891, SEQ ID No. 15 6845, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia pneumoniae cell wall anchored surface polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 267, SEQ ID No. 271, SEQ ID No. 419, SEQ ID No. 590, SEQ ID No. 932, SEQ ID No. 20 6844, SEQ ID No. 6847, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments not found in *Chlamydia trachomatis* (Blastp P>e<sup>-10</sup>), and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 7, SEQ ID No. 8, SEQ ID No. 9, SEQ ID No. 16, SEQ ID No. 17, SEQ ID No. 18, SEQ ID No. 19, SEQ ID No. 20, SEQ ID No. 21, SEQ ID No. 22, SEQ ID No. 1254, SEQ ID No. 23, SEQ ID No. 1255, SEQ ID No. 24, SEQ ID No. 1139, SEQ ID No. 1140, SEQ ID No. 46, SEQ ID No. 47, SEQ ID No. 51, SEQ ID No. 60, SEQ ID No. 1256, SEQ ID No. 61, SEQ ID No. 62, SEQ ID No. 63, SEQ ID No. 64, SEQ ID No. 1257, SEQ ID No. 65, SEQ ID No. 66, SEQ ID No. 67, SEQ ID No. 68,
SEQ ID No. 1143, SEQ ID No. 1145, SEQ ID No. 83, SEQ ID No. 84, SEQ ID No. 1146, SEQ ID No. 85, SEQ ID No. 86, SEQ ID No. 87, SEQ ID No. 1258, SEQ ID No. 116, SEQ ID No. 117, SEQ ID No. 125, SEQ ID No. 1148, SEQ ID No. 143, SEQ ID No. 1150, SEQ ID No. 1151, SEQ ID No. 144, SEQ ID No. 145, SEQ ID No. 147, SEQ ID No. 148, SEQ ID No. 149, SEQ ID No. 150, SEQ ID No. 167, SEQ ID No. 1261, SEQ ID No. 1156, SEQ ID No. 1157, SEQ ID No. 178, SEQ ID No. 179, SEQ ID No. 1158, SEQ ID No. 182, SEQ ID No. 183, SEQ ID No. 184, SEQ ID No. 185, SEQ ID No. 179, SEQ ID No. 1159, SEQ ID No. 186, SEQ ID No. 1160, SEQ ID No. 187, SEQ ID No. 188, SEQ ID No. 189, SEQ ID No. 189, SEQ ID No. 189, SEQ ID No. 188, SEQ ID No. 189, SEQ ID No. 180, SEQ ID No

No. 190, SEQ ID No. 1161, SEQ ID No. 1162, SEQ ID No. 191, SEQ ID No. 192, SEQ ID No. 194, SEQ ID No. 195, SEQ ID No. 1163, SEQ ID No. 196, SEQ ID No. 201, SEQ ID No. 202, SEQ ID No. 209, SEO ID No. 212, SEO ID No. 221, SEO ID No. 224, SEO ID No. 1167, SEO ID No. 226, SEO ID No. 227, SEQ ID No. 228, SEQ ID No. 229, SEQ ID No. 230, SEQ ID No. 231, SEQ ID No. 5 232, SEO ID No. 1169, SEQ ID No. 1170, SEQ ID No. 1171, SEQ ID No. 234, SEQ ID No. 235, SEQ ID No. 236, SEQ ID No. 1172, SEQ ID No. 243, SEQ ID No. 251, SEQ ID No. 252, SEQ ID No. 1176, SEQ ID No. 253, SEQ ID No. 255, SEQ ID No. 254, SEQ ID No. 256, SEQ ID No. 1177, SEQ ID No. 1178, SEQ ID No. 262, SEQ ID No. 263, SEQ ID No. 1264, SEQ ID No. 278, SEQ ID No. 279. SEO ID No. 1180, SEO ID No. 280, SEO ID No. 290, SEQ ID No. 291, SEQ ID No. 292, SEQ 10 ID No. 296, SEQ ID No. 1181, SEQ ID No. 297, SEQ ID No. 298, SEQ ID No. 300, SEQ ID No. 1265, SEO ID No. 322, SEQ ID No. 324, SEQ-ID No. 325, SEQ ID No. 370, SEQ ID No. 1186, SEQ ID No. 371, SEQ ID No. 372, SEQ ID No. 1187, SEQ ID No. 373, SEQ ID No. 378, SEQ ID No. 1266, SEQ ID No. 382, SEQ ID No. 383, SEQ ID No. 384, SEQ ID No. 385, SEQ ID No. 386, SEQ ID No. 1188, SEQ ID No. 1189, SEQ ID No. 391, SEQ ID No. 392, SEQ ID No. 398, SEQ ID No. 15 400, SEQ ID No. 403, SEQ ID No. 1191, SEQ ID No. 423, SEQ ID No. 435, SEQ ID No. 445, SEQ ID No. 450, SEQ ID No. 1193, SEQ ID No. 456, SEQ ID No. 460, SEQ ID No. 461, SEQ ID No. 465, SEO ID No. 1196, SEQ ID No. 471, SEQ ID No. 473, SEQ ID No. 475, SEQ ID No. 481, SEQ ID No. 484, SEQ ID No. 487, SEQ ID No. 488, SEQ ID No. 489, SEQ ID No. 490, SEQ ID No. 491, SEO ID No. 492, SEQ ID No. 493, SEQ ID No. 494, SEQ ID No. 495, SEQ ID No. 496, SEQ ID No. 20 497, SEQ ID No. 498, SEQ ID No. 499, SEQ ID No. 502, SEQ ID No. 1267, SEQ ID No. 1268, SEQ ID No. 508, SEQ ID No. 510, SEQ ID No. 509, SEQ ID No. 512, SEQ ID No. 515, SEQ ID No. 519, SEQ ID No. 1197, SEQ ID No. 521, SEQ ID No. 1198, SEQ ID No. 522, SEQ ID No. 524, SEQ ID No. 528, SEQ ID No. 534, SEQ ID No. 537, SEQ ID No. 1269, SEQ ID No. 1270, SEQ ID No. 548, SEO ID No. 551, SEO ID No. 557, SEQ ID No. 1201, SEQ ID No. 1203, SEQ ID No. 562, SEQ ID 25 No. 566, SEQ ID No. 593, SEQ ID No. 595, SEQ ID No. 600, SEQ ID No. 1271, SEQ ID No. 604, SEO ID No. 611, SEQ ID No. 612, SEQ ID No. 614, SEQ ID No. 616, SEQ ID No. 625, SEQ ID No. 627, SEO ID No. 628, SEQ ID No. 629, SEQ ID No. 631, SEQ ID No. 641, SEQ ID No. 1272, SEQ ID No. 648, SEQ ID No. 1212, SEQ ID No. 663, SEQ ID No. 685, SEQ ID No. 707, SEQ ID No. 714, SEO ID No. 715, SEQ ID No. 716, SEQ ID No. 717, SEQ ID No. 722, SEQ ID No. 746, SEQ ID No. 30 1273, SEQ ID No. 761, SEQ ID No. 764, SEQ ID No. 770, SEQ ID No. 1217, SEQ ID No. 783, SEQ ID No. 1274, SEO ID No. 803, SEQ ID No. 815, SEQ ID No. 1220, SEQ ID No. 835, SEQ ID No. 1221, SEQ ID No. 844, SEQ ID No. 845, SEQ ID No. 846, SEQ ID No. 847, SEQ ID No. 848, SEQ ID No. 849, SEQ ID No. 850, SEQ ID No. 851, SEQ ID No. 1275, SEQ ID No. 852, SEQ ID No. 862, SEQ ID No. 1276, SEQ ID No. 1277, SEQ ID No. 873, SEQ ID No. 1223, SEQ ID No. 892, SEQ ID 35 No. 919, SEQ ID No. 1225, SEQ ID No. 1278, SEQ ID No. 926, SEQ ID No. 1228, SEQ ID No. 1229, SEQ ID No. 1230, SEQ ID No. 1279, SEQ ID No. 1281, SEQ ID No. 1282, SEQ ID No. 1283, SEO ID No. 948, SEO ID No. 950, SEQ ID No. 949, SEQ ID No. 951, SEQ ID No. 980, SEQ ID No. 982, SEQ ID No. 1233, SEQ ID No. 999, SEQ ID No. 1000, SEQ ID No. 1001, SEQ ID No. 1002, SEQ ID No. 1008, SEQ ID No. 1285, SEQ ID No. 1235, SEQ ID No. 1016, SEQ ID No. 1019, SEQ ID No. 1027, SEQ ID No. 1036, SEQ ID No. 1241, SEQ ID No. 1048, SEQ ID No. 1049, SEQ ID No. 1050, SEQ ID No. 1053, SEQ ID No. 1054, SEQ ID No. 1064, SEQ ID No. 1076, SEQ ID No. 1091, SEQ ID No. 1288, SEQ ID No. 1093, SEQ ID No. 1289, SEQ ID No. 1101, SEQ ID No. 1103, SEQ ID No. 1245, SEQ ID No. 1246, SEQ ID No. 1247, SEQ ID No. 1290, SEQ ID No. 1291, SEQ ID No. 1115, SEQ ID No. 1116, SEQ ID No. 1118, SEQ ID No. 1120, SEQ ID No. 1249, SEQ ID No. 1121, SEQ ID No. 1250, SEQ ID No. 1126, SEQ ID No. 1251, SEQ ID No. 1127, SEQ ID No. 1128, SEQ ID No. 1130, SEQ ID No. 1129, SEQ ID No. 1131, SEQ ID No. 1136, SEQ ID No. 1253, SEQ ID No. 106844, SEQ ID No. 6846, SEQ ID No. 6847, SEQ ID No. 6848, and one of their representative fragments

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a Chlamydia pneumoniae polypeptide or one of its representative fragments which is involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of 15 cofactors, and in that it is chosen from the polypeptides having the following sequences: SEO ID No. 2; SEQ ID No. 55; SEQ ID No. 56; SEQ ID No. 69; SEQ ID No. 75; SEQ ID No. 80; SEO ID No. 100; SEO ID No. 110; SEQ ID No. 114; SEQ ID No. 120; SEQ ID No. 121; SEQ ID No. 157; SEQ ID No. 160; SEQ ID No. 161; SEQ ID No. 172; SEQ ID No. 180; SEQ ID No. 181; SEO ID No. 198; SEQ ID No. 200; SEQ ID No. 225; SEQ ID No. 248; SEQ ID No. 249; SEQ ID 20 No. 276; SEQ ID No. 277; SEQ ID No. 318; SEQ ID No. 319; SEQ ID No. 320; SEQ ID No. 323; SEQ ID No. 331; SEQ ID No. 347; SEQ ID No. 375; SEQ ID No. 376; SEQ ID No. 381; SEQ ID No. 393; SEO ID No. 394; SEO ID No. 395; SEO ID No. 396; SEQ ID No. 409; SEQ ID No. 446; SEO ID No. 447; SEO ID No. 448; SEO ID No. 449; SEO ID No. 513; SEO ID No. 516; SEO ID No. 571; SEQ ID No. 647; SEQ ID No. 662; SEQ ID No. 697; SEQ ID No. 718; SEQ ID No. 793; 25 SEO ID No. 794; SEO ID No. 808; SEO ID No. 809; SEO ID No. 838; SEO ID No. 839; SEO ID No. 840; SEQ ID No. 853; SEQ ID No. 854; SEQ ID No. 918; SEQ ID No. 923; SEQ ID No. 929; SEQ ID No. 931; SEQ ID No. 938; SEQ ID No. 939; SEQ ID No. 958; SEQ ID No. 959; SEQ ID No. 960; SEQ ID No. 966; SEQ ID No. 995; SEQ ID No. 1021; SEQ ID No. 1040; SEQ ID No. 1041; SEQ ID No. 1042; SEQ ID No. 1085; SEQ ID No. 1100; SEQ ID No. 1102; SEQ ID 30 No. 1117; SEQ ID No. 1118; SEQ ID No. 1119; SEQ ID No. 1120; SEQ ID No. 1135 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the intermediate metabolism of nucleotides or nucleic acids, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 77; SEQ ID No. 78; SEQ ID No. 138; SEQ ID No. 189; SEQ ID No. 190; SEQ ID No. 233; SEQ ID No. 246; SEQ ID No. 338; SEQ ID No. 412; SEQ ID No. 421; SEQ ID No. 438;

SEQ ID No. 607; SEQ ID No. 648; SEQ ID No. 657; SEQ ID No. 740; SEQ ID No. 783; SEQ ID No. 967; SEQ ID No. 989; SEQ ID No. 990; SEQ ID No. 992; SEQ ID No. 1011; SEQ ID No. 1058; SEQ ID No. 1059; SEQ ID No. 1073; SEQ ID No. 1074 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of nucleic acids, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 14; SEQ ID No. 59; SEQ ID No. 70; SEQ ID No. 71; SEQ ID No. 97; SEQ ID No. 113; SEQ ID No. 137; SEQ ID No. 141; SEQ ID No. 169; SEQ ID No. 285; SEQ ID No. 287; SEQ ID No. 288; SEQ ID No. 313; SEQ ID No. 326; SEQ ID No. 358; SEQ ID No. 411; SEQ ID No. 443; SEQ ID No. 548; SEQ ID No. 569; SEQ ID No. 601; SEQ ID No. 651; SEQ ID No. 654; SEQ ID No. 658; SEQ ID No. 659; SEQ ID No. 664; SEQ ID No. 665; SEQ ID No. 694; SEQ ID No. 698; SEQ ID No. 704; SEQ ID No. 760; SEQ ID No. 762; SEQ ID No. 763; SEQ ID No. 786; SEQ ID No. 787; SEQ ID No. 788; SEQ ID No. 801; SEQ ID No. 802; SEQ ID No. 812; SEQ ID No. 819; SEQ ID No. 822; SEQ ID No. 870; SEQ ID No. 897; SEQ ID No. 898; SEQ ID No. 908; SEQ ID No. 916; SEQ ID No. 954; SEQ ID No. 955; SEQ ID No. 961; SEQ ID No. 983; SEQ ID No. 996; SEQ ID No. 1007; SEQ ID No. 1012; SEQ ID No. 1013; SEQ ID No. 1014; SEQ ID No. 1015; SEQ ID No. 1038; SEQ ID No. 1137 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of amino acids or polypeptides, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 99; SEQ ID No. 111; SEQ ID No. 127; SEQ ID No. 134; SEQ ID No. 140; SEQ ID No. 174; SEQ ID No. 175; SEQ ID No. 176; SEQ ID No. 353; SEQ ID No. 377; SEQ ID No. 404; SEQ ID No. 523; SEQ ID No. 539; SEQ ID No. 559; SEQ ID No. 561; SEQ ID No. 586; SEQ ID No. 598; SEQ ID No. 609; SEQ ID No. 636; SEQ ID No. 687; SEQ ID No. 700; SEQ ID No. 701; SEQ ID No. 759; SEQ ID No. 790; SEQ ID No. 857; SEQ ID No. 861; SEQ ID No. 904; SEQ ID No. 936; SEQ ID No. 952; SEQ ID No. 962; SEQ ID No. 963; SEQ ID No. 964; SEQ ID No. 965; SEQ ID No. 991; SEQ ID No. 1003; SEQ ID No. 1004; SEQ ID No. 1005; SEQ ID No. 1018; SEQ ID No. 1067; SEQ ID No. 1110; SEQ ID No. 1111; SEQ ID No. 1112; SEQ ID No. 1114; SEQ ID No. 1121; SEQ ID No. 1122; SEQ ID No. 1123; SEQ ID No. 1124; SEQ ID No. 1125 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of polypeptides, and in that it is chosen from the polypeptides

having the following sequences:

SEQ ID No. 4; SEQ ID No. 44; SEQ ID No. 45; SEQ ID No. 48; SEQ ID No. 54; SEQ ID No. 112; SEQ ID No. 130; SEQ ID No. 155; SEQ ID No. 163; SEQ ID No. 212; SEQ ID No. 257; SEQ ID No. 307; SEQ ID No. 343; SEQ ID No. 405; SEQ ID No. 416; SEQ ID No. 458; SEQ ID No. 540; SEQ ID No. 541; SEQ ID No. 542; SEQ ID No. 543; SEQ ID No. 544; SEQ ID No. 560; SEQ ID No. 594; SEQ ID No. 652; SEQ ID No. 699; SEQ ID No. 723; SEQ ID No. 747; SEQ ID No. 817; SEQ ID No. 827; SEQ ID No. 871; SEQ ID No. 909; SEQ ID No. 910; SEQ ID No. 911; SEQ ID No. 912; SEQ ID No. 1023; SEQ ID No. 1051; SEQ ID No. 1052; SEQ ID No. 1081 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the metabolism of fatty acids, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 76; SEQ ID No. 284; SEQ ID No. 308; SEQ ID No. 309; SEQ ID No. 310; SEQ ID No. 311; SEQ ID No. 312; SEQ ID No. 425; SEQ ID No. 433; SEQ ID No. 565; SEQ ID No. 688; SEQ ID No. 690; SEQ ID No. 691; SEQ ID No. 767; SEQ ID No. 797; SEQ ID No. 894; SEQ ID No. 895; SEQ ID No. 994; SEQ ID No. 1020; SEQ ID No. 1030; SEQ ID No. 1033; SEQ ID No. 1034; SEQ ID No. 1046; SEQ ID No. 1047; SEQ ID No. 1057 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the synthesis of the wall, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 49; SEQ ID No. 50; SEQ ID No. 177; SEQ ID No. 178; SEQ ID No. 245; SEQ ID
 No. 610; SEQ ID No. 972; SEQ ID No. 974; SEQ ID No. 978; SEQ ID No. 1037 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the transcription, translation and/or maturation process, and in that it is chosen 30 from the polypeptides having the following sequences:

SEQ ID No. 90; SEQ ID No. 92; SEQ ID No. 131; SEQ ID No. 151; SEQ ID No. 199; SEQ ID No. 333; SEQ ID No. 334; SEQ ID No. 336; SEQ ID No. 379; SEQ ID No. 589; SEQ ID No. 590; SEQ ID No. 619; SEQ ID No. 630; SEQ ID No. 649; SEQ ID No. 739; SEQ ID No. 741; SEQ ID No. 806; SEQ ID No. 821; SEQ ID No. 843; SEQ ID No. 968; SEQ ID No. 971; SEQ ID No. 1061 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* ribosomal polypeptide or one of its representative

fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 93; SEQ ID No. 94; SEQ ID No. 95; SEQ ID No. 136; SEQ ID No. 259; SEQ ID No. 332; SEQ ID No. 348; SEQ ID No. 583; SEQ ID No. 584; SEQ ID No. 588; SEQ ID No. 591; SEQ ID No. 592; SEQ ID No. 663; SEQ ID No. 666; SEQ ID No. 667; SEQ ID No. 669; SEQ ID No. 670; SEQ ID No. 671; SEQ ID No. 672; SEQ ID No. 673; SEQ ID No. 674; SEQ ID No. 675; SEQ ID No. 676; SEQ ID No. 677; SEQ ID No. 678; SEQ ID No. 679; SEQ ID No. 680; SEQ ID No. 681; SEQ ID No. 683; SEQ ID No. 684; SEQ ID No. 738; SEQ ID No. 781; SEQ ID No. 1008; SEO ID No. 1024; SEO ID No. 1025; SEQ ID No. 1066 and one of their representative fragments.

Preferably, the invention also relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* transport polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 40; SEQ ID No. 41; SEQ ID No. 52; SEQ ID No. 105; SEQ ID No. 106; SEQ ID No. 107; SEQ ID No. 109; SEQ ID No. 133; SEQ ID No. 210; SEQ ID No. 211; SEQ ID No. 214; SEQ ID No. 215; SEQ ID No. 216; SEQ ID No. 217; SEQ ID No. 218; SEQ ID No. 219; SEQ ID No. 220; SEQ ID No. 223; SEQ ID No. 242; SEQ ID No. 260; SEQ ID No. 293; SEQ ID No. 299; SEQ ID No. 366; SEQ ID No. 369; SEQ ID No. 575; SEQ ID No. 602; SEQ ID No. 638; SEQ ID No. 639; SEQ ID No. 640; SEQ ID No. 643; SEQ ID No. 653; SEQ ID No. 702; SEQ ID No. 703; SEQ ID No. 724; SEQ ID No. 732; SEQ ID No. 855; SEQ ID No. 856; SEQ ID No. 901; SEQ ID No. 906; SEQ ID No. 933; SEQ ID No. 942; SEQ ID No. 1043; SEQ ID No. 1086; SEQ ID No. 1086; SEQ ID No. 1105 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the virulence process, and in that it is chosen from the polypeptides having the following sequences:

25 SEQ ID No. 546; SEQ ID No. 550; SEQ ID No. 778; SEQ ID No. 779; SEQ ID No. 886 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia pneumoniae* polypeptide or one of its representative fragments which is involved in the secretory system and/or which is secreted, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 751; SEQ ID No. 874; SEQ ID No. 875; SEQ ID No. 876; SEQ ID No. 883; SEQ ID No. 884; SEQ ID No. 885 and one of their representative fragments.

The secreted polypeptides, including the Type III and other, non-Type III secreted polypeptides, of the present invention, as well as the corresponding nucleotide sequences, may be detected by techniques known to persons skilled in the art, such as for example the techniques using cloning combined with vectors allowing the expression of the said polypeptides fused to export markers such as the *luc* gene for luciferase or the *PhoA* gene for alkaline phosphatase.

Preferably, the invention relates to a polypeptide according characterized in that it is a polypeptide specific to Chlamydia pneumoniae or one of its representative fragments (with a Blast E value of >10-5), and in that it is chosen from the polypeptides having the following sequences:

5 SEQ ID No. 7; SEQ ID No. 8; SEQ ID No. 17; SEQ ID No. 18; SEQ ID No. 19; SEQ ID No. 20: SEO ID No. 22; SEQ ID No. 23; SEQ ID No. 24; SEQ ID No. 51; SEQ ID No. 60; SEQ ID No. 63; SEQ ID No. 65; SEQ ID No. 66; SEQ ID No. 67; SEQ ID No. 83; SEQ ID No. 84; SEO ID No. 86; SEQ ID No. 87; SEQ ID No. 125; SEQ ID No. 143; SEQ ID No. 144; SEQ ID No. 179; SEO ID No. 182; SEO ID No. 184; SEQ ID No. 185; SEQ ID No. 187; SEQ ID No. 221; 10 SEQ ID No. 252; SEQ ID No. 254;; SEQ ID No. 278; SEQ ID No. 279; SEQ ID No. 387; SEQ ID No. 388; SEQ-ID No. 397; -SEQ-ID No. 1048; SEQ-ID No. 1049; SEQ-ID No. 1050; SEQ-ID No. 1128; SEQ ID No. 1130; SEQ ID No. 1131 and one of their representative fragments.

In general, in the present invention, the functional group to which a polypeptide of the invention belongs, as well as its corresponding nucleotide sequence, may be determined either by 15 comparative analogy with sequences already known, or by the use of standard techniques of biochemistry, of cytology combined with the techniques of genetic engineering such as immunoaffinity, localization by immunolabelling, differential extraction, measurement of enzymatic activity, study of the activity inducing or repressing expression or the study of expression in E. coli.

It is clearly understood, on the one hand, that, in the present invention, the nucleotide 20 sequences (ORF) and the amino acid sequences (SEQ ID No. 2 to SEQ ID No. 1291 and SEQ ID No. 6844 to SEQ ID No. 6848) which are listed by functional group, are not exhaustive within the group considered. Moreover, it is also clearly understood that, in the present invention, a nucleotide sequence (ORF) or an amino acid sequence mentioned within a given functional group may also be part of another group taking into account, for example, the interrelationship between the groups listed. 25 Accordingly, and as an example of this interrelationship, an exported and/or secreted polypeptide as well as its coding nucleotide sequence may also be involved in the Chlamydia pneumoniae virulence process by modifying the defense mechanism of the infected host cell, or a transmembrane polypeptide or its coding nucleotide sequence is also part of the polypeptides or coding nucleotide sequences of the cellular envelope.

The subject of the present invention is also the nucleotide and/or polypeptide sequences according to the invention, characterized in that the said sequences are recorded on a medium, called recording medium, whose type and nature facilitate the reading, the analysis and the exploitation of the said sequences. These media may of course also contain other information extracted from the present invention, such as in particular the analogies with already known sequences, such as those 35 mentioned in Table 1 of the present description, and/or may contain, in addition, information relating to the nucleotide and/or polypeptide sequences of other microorganisms so as to facilitate the comparative analysis and the exploitation of the results obtained.

Among these recording media, computer-readable media, such as magnetic, optical, electrical and hybrid media such as, for example, floppy disks, CD-ROMs or recording cassettes, are preferred in particular.

The invention also relates to nucleotide sequences which can be used as primer or probe, characterized in that the said sequences are chosen from the nucleotide sequences according to the invention.

The invention relates, in addition, to the use of a nucleotide sequence according to the invention, as primer or probe, for the detection and/or amplification of nucleic acid sequences.

The nucleotide sequences according to the invention may thus be used to amplify nucleotide sequences, in particular by the PCR technique (polymerase chain reaction) (Erlich, 1989; Innis et al., 1990; Rolfs et al., 1991, and White et al., 1997).

These oligodeoxyribonucleotide or oligoribonucleotide primers correspond to representative nucleotide fragments, and are advantageously at least 8 nucleotides, preferably at least 12 nucleotides, 15 nucleotides and still more preferably at least 20 nucleotides long.

Other techniques for amplifying the target nucleic acid may be advantageously used as alternatives to PCR.

The nucleotide sequences of the invention, in particular the primers according to the invention, may also be used in other methods for amplifying a target nucleic acid, such as:

- the TAS (Transcription-based Amplification System) technique described by Kwoh et al. in 1989;
- 20 the 3SR (Self-Sustained Sequence Replication) technique described by Guatelli et al. in 1990;
  - the NASBA (Nucleic Acid Sequence Based Amplification) technique described by Kievitis et al. in 1991;
  - the SDA (Strand Displacement Amplification) technique (Walker et al., 1992);
  - the TMA (Transcription Mediated Amplification) technique.
- The polynucleotides of the invention may also be used in techniques for amplifying or for modifying the nucleic acid serving as probe, such as:
  - the LCR (Ligase Chain Reaction) technique described by Landegren et al. in 1988 and perfected by Barany et al. in 1991, which uses a thermostable ligase;
  - the RCR (Repair Chain Reaction) technique described by Segev in 1992;
- 30 the CPR (Cycling Probe Reaction) technique described by Duck et al. in 1990;
  - the Q-beta-replicase amplification technique described by Miele et al. in 1983 and perfected in particular by Chu et al. in 1986, Lizardi et al. in 1988, and then by Burg et al. as well as by Stone et al. in 1996.

The invention also relates to the nucleotide sequences of fragments which can be
35 obtained by amplification with the aid of at least one primer according to the invention. The present
invention encompasses both hybridization probes and primers. In general, the complementary probes
should be of a length sufficient to form a stable hybrid complex with the target sequences. Primers,

while complementary to the target sequences need not form stable hybridization complexes with the target sequences alone. Rather, primers form stable complexes with the target sequences in the presence of polymerase to permit extension of the primer.

In the case where the target polynucleotide to be detected is possibly an RNA, for example an mRNA, it will be possible to use, prior to the use of an amplification reaction with the aid of at least one primer according to the invention or to the use of a method of detection with the aid of at least one probe of the invention, a reverse transcriptase-type enzyme so as to obtain a cDNA from the RNA contained in the biological sample. The cDNA obtained will then serve as target for the primer(s) or the probe(s) used in the amplification or detection method according to the invention.

The detection probe will be chosen so that it hybridizes with the target sequence or the amplicon generated from the target sequence. Such a detection probe will advantageously have as sequence a sequence of at least 12 nucleotides, in particular of at least 20 nucleotides, and preferably at least 100 nucleotides.

The invention also comprises the nucleotide sequences which can be used as probe or primer according to the invention, characterized in that they are labelled with a radioactive compound or with a nonradioactive compound.

The nonlabelled nucleotide sequences may be used directly as probes or primers; however, the sequences are generally labelled with a radioactive element (<sup>32</sup>P, <sup>35</sup>S, <sup>3</sup>H, <sup>125</sup>I) or with a nonradioactive molecule (biotin, acetylaminofluorene, digoxigenin, 5-bromo-deoxyuridine, 20 fluorescein) so as to obtain probes which can be used in numerous applications.

Examples of nonradioactive labelling of nucleotide sequences are described, for example, in French patent No. 78,10975 or by Urdea et al. or by Sanchez-Pescador et al. in 1988.

In the latter case, one of the labelling methods described in patents FR-2 422 956 and FR-2 518 755 may also be used.

The invention also relates to the nucleotide sequences of fragments which can be obtained by hybridization with the aid of at least one probe according to the invention.

The hybridization technique may be performed in various ways (Matthews et al., 1988).

The most common method consists in immobilizing the nucleic acid extracted from Chlamydia pneumoniae cells on a support (such as nitrocellulose, nylon, polystyrene) and in incubating, under well-defined conditions, the target nucleic acid immobilized with the probe. After hybridization, the excess probe is removed and the hybrid molecules formed are detected by the appropriate method (measurement of the radioactivity, of the fluorescence or of the enzymatic activity linked to the probe).

The invention also comprises the nucleotide sequences according to the invention, 35 characterized in that they are covalently or noncovalently immobilized on a support.

According to another advantageous embodiment of the nucleic sequences according to the invention, the latter may be used immobilized on a support and may thus serve to capture, through

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specific hybridization, the target nucleic acid obtained from the biological sample to be tested. If necessary, the solid support is separated from the sample and the hybridization complex formed between the so-called capture probe and the target nucleic acid is then detected by means of a second probe, called detection probe, labelled with an easily detectable element.

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The nucleotide sequences according to the invention may also be used in new analytical systems, DNA chips, which allow sequencing, the study of mutations and of the expression of genes, and which are currently of interest given their very small size and their high capacity in terms of number of analyses.

The principle of the operation of these chips is based on molecular probes, most often oligonucleotides, which are attached onto a miniaturized surface, generally of the order of a few square centimetres. During an analysis, a sample containing fragments of a target nucleic acid to be analysed, for example DNA or RNA labelled, for example, after amplification, is deposited onto the DNA chip in which the support has been coated beforehand with probes. Bringing the labelled target sequences into contact with the probes leads to the formation, through hybridization, of a duplex according to the rule of pairing defined by J.D. Watson and F. Crick. After a washing step, analysis of the surface of the chip allows the effective hybridizations to be located by means of the signals emitted by the labels tagging the target. A hybridization fingerprint results from this analysis which, by appropriate computer processing, will make it possible to determine information such as the presence of specific fragments in the sample, the determination of sequences and the presence of mutations.

The chip consists of a multitude of molecular probes, precisely organized or arrayed on a solid support whose surface is miniaturized. It is at the centre of a system where other elements (imaging system, microcomputer) allow the acquisition and interpretation of a hybridization fingerprint.

The hybridization supports are provided in the form of flat or porous surfaces (pierced with wells) composed of various materials. The choice of a support is determined by its physicochemical properties, or more precisely, by the relationship between the latter and the conditions under which the support will be placed during the synthesis or the attachment of the probes or during the use of the chip. It is therefore necessary, before considering the use of a particular support (R.S. Matson et al., 1994), to consider characteristics such as its stability to pH, its physical strength, its reactivity and its chemical stability as well as its capacity to nonspecifically bind nucleic acids. Materials such as glass, silicon and polymers are commonly used. Their surface is, in a first step, called "functionalization", made reactive towards the groups which it is desired to attach thereon. After the functionalization, so-called spacer molecules are grafted onto the activated surface. Used as intermediates between the surface and the probe, these molecules of variable size render unimportant the surface properties of the supports, which often prove to be problematic for the synthesis or the attachment of the probes and for the hybridization.

Among the hybridization supports, there may be mentioned glass which is used, for

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example, in the method of in situ synthesis of oligonucleotides by photochemical addressing developed by the company Affymetrix (E.L. Sheldon, 1993), the glass surface being activated by silane. Genosensor Consortium (P. Mérel, 1994) also uses glass slides carrying wells 3 mm apart, this support being activated with epoxysilane.

Polymers or silicon may also be mentioned among these hybridization supports. For example, the Andrein Mirzabekov team has developed a chip consisting of polyacrylamide squares polymerized on a silanized glass surface (G. Yershov et al., 1996). Several teams use silicon, in particular the IFOS laboratory of Ecole Centrale of Lyon which uses a silicon semiconductor substrate which is p-doped by introducing it into its crystalline structure atoms whose valency is different from 10 that of silicon. Various types of metals, in particular gold and platinum, may also be used as support (Genosensor Consortium (K. Beattie et al., 1993)).

The probes according to the invention may be synthesized directly in situ on the supports of the DNA chips. This in situ synthesis may be carried out by photochemical addressing (developed by the company Affymax (Amsterdam, Holland) and exploited industrially by its subsidiary 15 Affymetrix (United States)) or based on the VLSIPS (very large scale immobilized polymer synthesis) technology (S.P.A. Fodor et al., 1991) which is based on a method of photochemically directed combinatory synthesis and the principle of which combines solid-phase chemistry, the use of photolabile protecting groups and photolithography.

The probes according to the invention may be attached to the DNA chips in various ways 20 such as electrochemical addressing, automated addressing or the use of probe printers (T. Livache et al., 1994; G. Yershov et al., 1996; J. Derisi et al., 1996, and S. Borman, 1996).

The revealing of the hybridization between the probes of the invention, deposited or synthesized in situ on the supports of the DNA chips, and the sample to be analysed, may be determined, for example, by measurement of fluorescent signals, by radioactive counting or by 25 electronic detection.

The use of fluorescent molecules such as fluorescein constitutes the most common method of labelling the samples. It allows direct or indirect revealing of the hybridization and allows the use of various fluorochromes.

Affymetrix currently provides an apparatus or a scanner designed to read its Gene Chip™ 30 chips. It makes it possible to detect the hybridizations by scanning the surface of the chip in confocal microscopy (R.J. Lipshutz et al., 1995). Other methods of detecting fluorescent signals have been tested: coupling of an epifluorescence microscope and a CCD camera (G. Yershov et al., 1996), the use of an optical fibre collecting system (E.L. Sheldon, 1993). A conventional method consists in carrying out an end labelling, with phosphorus 32, of the target sequences, by means of an appropriate 35 apparatus, the Phosphorimager (marketed by Molecular Dynamics). The electronic detection is based on the principle that the hybridization of two nucleic acid molecules is accompanied by physical phenomena which can be quantified under certain conditions (system developed by Ecole Centrale of

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Lyon and called GEN-FET (GEN field effect transistor)). Genosensor Consortium and the company Beckman Instruments who are developing an electronic chip or Permittivity Chips<sup>TM</sup> may also be mentioned (K. Beattie et al., 1993).

The nucleotide sequences according to the invention may thus be used in DNA chips to carry out the analysis of mutations. This analysis is based on the production of chips capable of analysing each base of a nucleotide sequence according to the invention.

The nucleotide sequences according to the invention may also be used in DNA chips to carry out the analysis of the expression of the Chlamydia pneumoniae genes. This analysis of the expression of Chlamydia pneumoniae genes is based on the use of chips where probes of the invention, chosen for their specificity to characterize a given gene, are present (D.J. Lockhart et al., 1996; D.D. Shoemaker et al., 1996). For the methods of analysis of gene expression using the DNA chips, reference may, for example, be made to the methods described by D.J. Lockhart et al. (1996) and Sosnowsky et al. (1997) for the synthesis of probes in situ or for the addressing and the attachment of previously synthesized probes. The target sequences to be analysed are labelled and in general fragmented into sequences of about 50 to 100 nucleotides before being hybridized onto the chip. After washing as described, for example, by D.J. Lockhart et al. (1996) and application of different electric fields (Sosnowsky et al., 1997), the labelled compounds are detected and quantified, the hybridizations being carried out at least in duplicate. Comparative analyses of the signal intensities obtained with respect to the same probe for different samples and/or for different probes with the same sample, determine the differential expression of RNA or of DNA derived from the sample.

The nucleotide sequences according to the invention may, in addition, be used in DNA chips where other nucleotide probes specific for other microorganisms are also present, and may allow the carrying out of a serial test allowing rapid identification of the presence of a microorganism in a sample.

Accordingly, the subject of the invention is also the nucleotide sequences according to the invention, characterized in that they are immobilized on a support of a DNA chip.

The DNA chips, characterized in that they contain at least one nucleotide sequence according to the invention, immobilized on the support of the said chip, also form part of the invention.

The said chips will preferably contain several probes or nucleotide sequences of the invention of different length and/or corresponding to different genes so as to identify, with greater certainty, the specificity of the target sequences or the desired mutation in the sample to be analysed.

Accordingly, the analyses carried out by means of primers and/or probes according to the invention, immobilized on supports such as DNA chips, will make it possible, for example, to identify, in samples, mutations linked to variations such as intraspecies variations. These variations may be correlated or associated with pathologies specific to the variant identified and will make it possible to select the appropriate treatment.

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The invention thus comprises a DNA chip according to the invention, characterized in that it contains, in addition, at least one nucleotide sequence of a microorganism different from Chlamydia pneumoniae, immobilized on the support of the said chip; preferably, the different microorganism will be chosen from an associated microorganism, a bacterium of the 5 Chlamydia family, and a variant of the species Chlamydia pneumoniae.

Another subject of the present invention is a vector for the cloning and/or the expression of a sequence, characterized in that it contains a nucleotide sequence according to the invention. Among the said vectors according to the invention, the vectors containing a nucleotide sequence encoding a polypeptide of the cellular, preferably outer, envelope of Chlamydia pneumoniae or one of 10 its representative fragments, are preferred. In a specific embodiment, the vectors contain a nucleotide sequence encoding a Chlamydia pneumoniae secreted polypeptide or one of its representative fragments or encoding a transport polypeptide, a surface exposed polypeptide, a lipoprotein or one of its representative fragments, a polypeptide involved in lipopolysaccharide (LPS) biosynthesis, a Type III and non-Type III secreted polypeptide, a polypeptide containing RGD attachment sites, a cell wall 15 anchored surface polypeptide, a polypeptide not found in Chlamydia trachomatis, a ribosomal polypeptide or a polypeptide involved in secretion, transcription, translation, maturation of proteins, a polypeptide involved in the synthesis of the wall, a polypeptide involved in the virulence, a polypeptide involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, a polypeptide involved in the metabolism of nucleotides, of amino acids, of nucleic acids 20 or of fatty acids of Chlamydia pneumoniae or one of their representative fragments, or a polypeptide specific to Chlamydia pneumoniae.

According to the invention, the vectors comprise the elements necessary to allow the expression and/or the secretion of the said nucleotide sequences in a given host cell, and form part of the invention. The vector should, in this case, comprise a promoter, signals for initiation and for termination of translation, as well as appropriate regions for regulation of transcription. It should be capable of being stably maintained in the host cell and may optionally possess particular signals specifying the secretion of the translated protein. These different elements are chosen according to the host cell used. To this effect, the nucleotide sequences according to the invention may be inserted into autonomously-replicating vectors within the chosen host, or integrative vectors in the chosen host.

Any of the standard methods known to those skilled in the art for the insertion of DNA fragments into a vector may be used to construct expression vectors containing a chimeric gene consisting of appropriate transcriptional/translational control signals and the protein coding sequences. These methods may include *in vitro* recombinant DNA and synthetic techniques and *in vivo* recombinants (genetic recombination).

Expression of a polypeptide, peptide or derivative, or analogs thereof encoded by a polynucleotide sequence in SEQ ID No. 1 or ORFs contained within SEQ ID No. 1 may be regulated by a second nucleic acid sequence so that the protein or peptide is expressed in a host transformed

For example, expression of a protein or peptide may with the recombinant DNA molecule. be controlled by any promoter/enhancer element known in the art. Promoters which may be used to control expression include, but are not limited to, the CMV promoter, the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 3' long terminal 5 repeat of Rous sarcoma virus (Yamamoto, et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al., 1982, Nature 296:39-42); prokaryotic expression vectors such as the 3-lactamase promoter (Villa-Kamaroff, et al., 1978, Proc. Natl. Acad. Sci. U.S.A. 75:3727-3731), or the tac promoter (DeBoer, et al., 1983, Proc. Natl. Acad. Sci. U.S.A. 80:21-25); see 10 also "Useful proteins from recombinant bacteria" in Scientific American, 1980, 242:74-94; plant expression vectors comprising the nopaline synthetase promoter region (Herrera-Estrella et al., 1983, Nature 303:209-213) or the cauliflower mosaic virus 35S RNA promoter (Gardner, et al., 1981, Nucl. Acids Res. 9:2871), and the promoter of the photosynthetic enzyme ribulose biphosphate carboxylase (Herrera-Estrella et al., 1984, Nature 310:115-120); promoter elements from yeast or other fungi such 15 as the Gal 4 promoter, the ADC (alcohol dehydrogenase) promoter, PGK (phosphoglycerol kinase) promoter, alkaline phosphatase promoter, and the following animal transcriptional control regions, which exhibit tissue specificity and have been utilized in transgenic animals: elastase I gene control region which is active in pancreatic acinar cells (Swift et al., 1984, Cell 38:639-646; Ornitz et al., 1986, Cold Spring Harbor Symp. Quant. Biol. <u>50</u>:399-409; MacDonald, 1987, Hepatology <u>7</u>:425-515); 20 insulin gene control region which is active in pancreatic beta cells (Hanahan, 1985, Nature 315:115-122), immunoglobulin gene control region which is active in lymphoid cells (Grosschedl et al., 1984, Cell 38:647-658; Adames et al., 1985, Nature 318:533-538; Alexander et al., 1987, Mol. Cell. Biol. 7:1436-1444), mouse mammary tumor virus control region which is active in testicular, breast, lymphoid and mast cells (Leder et al., 1986, Cell 45:485-495), albumin gene control region which is 25 active in liver (Pinkert et al., 1987, Genes and Devel. 1:268-276), alpha-fetoprotein gene control region which is active in liver (Krumlauf et al., 1985, Mol. Cell. Biol. 5:1639-1648; Hammer et al., 1987. Science 235:53-58; alpha 1-antitrypsin gene control region which is active in the liver (Kelsey et al., 1987, Genes and Devel. 1:161-171), beta-globin gene control region which is active in myeloid cells (Mogram et al., 1985, Nature 315:338-340; Kollias et al., 1986, Cell 46:89-94; myelin basic 30 protein gene control region which is active in oligodendrocyte cells in the brain (Readhead et al., 1987, Cell 48:703-712); myosin light chain-2 gene control region which is active in skeletal muscle (Sani, 1985, Nature 314:283-286), and gonadotropic releasing hormone gene control region which is active in the hypothalamus (Mason et al., 1986, Science 234:1372-1378).

The vectors according to the invention are, for example, vectors of plasmid or viral origin. In a specific embodiment, a vector is used that comprises a promoter operably linked to a protein or peptide-encoding a nucleic acid sequence in SEQ ID No. 1, or ORFs contained within SEQ ID No. 1, one or more origins of replication, and, optionally, one or more selectable markers (e.g., an

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antibiotic resistance gene). Expression vectors comprise regulatory sequences that control gene expression, including gene expression in a desired host cell. Preferred vectors for the expression of the polypeptides of the invention include the pET-type plasmid vectors (Promega) or pBAD plasmid vectors (Invitrogen). Furthermore, the vectors according to the invention are useful for transforming host cells so as to clone or express the nucleotide sequences of the invention.

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Expression can also be achieved using targeted homologous recombination to activate Chlamydia pneumoniae genes present in the cloned genomic DNA. A heterologous regulatory element may be inserted into a stable cell line or cloned microorganism, such that it is operatively linked with an endogenous Chlamydia pneumoniae gene present in the cloned genome, using techniques, such as targeted homologous recombination, which are well known to those of skill in the art (See, e.g., Chappel, U.S. Patent No. 4,215,051 and Skoultchi, WO 91/06667 each of which is incorporated herein in its entirety).

Expression vector/host cell systems containing inserts of polynucleotide sequences in SEQ ID No. 1 or ORFs within SEQ ID No. 1, which encode polypeptides, peptides or derivatives, or 15 analogs thereof, can be identified by three general approaches: (a) nucleic acid hybridization, (b) presence or absence of "marker" gene functions, and (c) expression of inserted sequences. In the first approach, the presence of a polynucleotide sequence inserted in an expression vector can be detected by nucleic acid hybridization using probes comprising sequences that are homologous to an inserted polynucleotide sequence. In the second approach, the recombinant vector/host system can be 20 identified and selected based upon the presence or absence of certain "marker" gene functions (e.g., thymidine kinase activity, resistance to antibiotics, transformation phenotype, occlusion body formation in baculovirus, etc.) caused by the insertion of a polynucleotide sequence in the vector. For example, if the polynucleotide sequence in SEQ ID No. 1 or ORFs within SEQ ID No. 1 is inserted within the marker gene sequence of the vector, recombinants containing the insert can be identified by 25 the absence of the marker gene function. In the third approach, recombinant expression vectors can be identified by assaying the product of the polynucleotide sequence expressed by the recombinant. Such assays can be based, for example, on the physical or functional properties of the expressed polypeptide in in vitro assay systems, e.g., binding with antibody, promotion of cell proliferation.

Once a particular recombinant DNA molecule is identified and isolated, several methods 30 known in the art may be used to propagate it. The clones identified may be introduced into an appropriate host cell by standard methods, such as for example lipofection, electroporation, and heat shock. Once a suitable host system and growth conditions are established, recombinant expression vectors can be propagated and prepared in quantity.

The invention also encompasses the host cells transformed by a vector according to the invention. These cells may be obtained by introducing into host cells a nucleotide sequence inserted into a vector as defined above, and then culturing the said cells under conditions allowing the replication and/or the expression of the transfected nucleotide sequence.

The host cell may be chosen from eukaryotic or prokaryotic systems, such as for example bacterial cells (Olins and Lee, 1993), but also yeast cells (Buckholz, 1993), as well as animal cells, in particular cultures of mammalian cells (Edwards and Aruffo, 1993), and in particular Chinese hamster ovary (CHO) cells, but also insect cells in which methods using baculoviruses for example may be used (Luckow, 1993).

Furthermore, a host cell strain may be chosen which modulates the expression of the inserted sequences, or modifies and processes the gene product in the specific fashion desired. Expression from certain promoters can be elevated in the presence of certain inducers; thus, expression of the genetically engineered polypeptide may be controlled. Furthermore, different host cells have characteristic and specific mechanisms for the translational and post-translational processing and modification (e.g., glycosylation, phosphorylation) of proteins. Appropriate cell-lines or host systems can be chosen to ensure the desired modification and processing of the foreign protein expressed. For example, expression in a bacterial system can be used to produce an unglycosylated core protein product. Expression in yeast will produce a glycosylated product. Expression in mammalian cells can be used to ensure "native" glycosylation of a heterologous protein. Furthermore, different vector/host expression systems may effect processing reactions to different extents.

A preferred host cell for the expression of the proteins of the invention consists of prokaryotic cells, such as Gram bacteria. A further preferred host cell according to the invention is a bacterium belonging to the *Chlamydia* family, more preferably belonging to the species *Chlamydia* pneumoniae or chosen from a microorganism associated with the species *Chlamydia pneumoniae*.

In other specific embodiments, the polypeptides, peptides or derivatives, or analogs thereof may be expressed as a fusion, or chimeric protein product (comprising the protein, fragment, analog, or derivative joined via a peptide bond to a heterologous protein sequence (of a different protein)). Such a chimeric product can be made by ligating the appropriate nucleic acid sequences encoding the desired amino acid sequences to each other by methods known in the art, in the proper coding frame, and expressing the chimeric product by methods commonly known in the art. Alternatively, such a chimeric product may be made by protein synthetic techniques, e.g., by use of a peptide synthesizer.

Genomic sequences can be cloned and expressed as translational gene products (i.e., 30 peptides, polypeptides, and proteins) or transcriptional gene products (i.e., antisense and ribozymes).

The invention further relates to the intracellular production of an antisense nucleic acid sequence of SEQ ID No. 1 by transcription from an exogenous sequence. For example, a vector can be introduced *in vivo* such that it is taken up by a cell, within which cell the vector or a portion thereof is transcribed, producing an antisense nucleic acid (RNA) of the invention. Such a vector would contain a sequence encoding an antisense nucleic acid. Such a vector can remain episomal or become chromosomally integrated, as long as it can be transcribed to produce the desired antisense RNA. Such vectors can be constructed by recombinant DNA technology methods standard in the art.

Vectors can be plasmid, viral, or others known in the art, used for replication and expression in mammalian cells. Expression of the sequence encoding the an antisense RNA can be by any promoter known in the art to act in mammalian, preferably human, cells. Such promoters can be inducible or constitutive. Such promoters include but are not limited to: the CMV promoter, the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 3N long terminal repeat of Rous sarcoma virus (Yamamoto et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al., 1982, Nature 296:39-42), etc.

In a specific embodiment, the antisense oligonucleotide comprises catalytic RNA, or a ribozyme (see, e.g., PCT International Publication WO 90/11364, published October 4, 1990; Sarver et al., 1990, Science 247:1222-1225). In another embodiment, the oligonucleotide is a 2N-0-methylribonucleotide (Inoue et al., 1987, Nucl. Acids Res. 15:6131-6148), or a chimeric RNA-DNA analog (Inoue et al., 1987, FEBS Lett. 215:327-330).

In another embodiment, the antisense nucleic acids of the invention comprise a sequence complementary to at least a portion of an RNA transcript of a polynucleotide sequence in SEQ ID No.

1. However, absolute complementarity, although preferred, is not required. A sequence "complementary to at least a portion of an RNA," as referred to herein, means a sequence having sufficient complementarity to be able to hybridize with the RNA, forming a stable duplex; in the case of double-stranded antisense nucleic acid sequence, a single strand of the duplex DNA may thus be tested, or triplex formation may be assayed. The ability to hybridize will depend on both the degree of complementarity and the length of the antisense nucleic acid. Generally, the longer the hybridizing nucleic acid, the more base mismatches with an RNA transcribed from SEQ ID No. 1 may contain and still form a stable duplex (or triplex, as the case may be). One skilled in the art can ascertain a tolerable degree of mismatch by use of standard procedures to determine the melting point of the hybridized complex.

The invention also relates to the animals, except humans, comprising one of the above-described transformed cells according to the invention.

The production of transgenic animals according to the invention overexpressing one or more of the *Chlamydia pneumoniae* genes will be preferably carried out on rats, mice or rabbits according to methods well known to persons skilled in the art such as viral or nonviral transfections. The transgenic animals overexpressing one or more of the said genes may be obtained by transfection of multiple copies of the said genes under the control of a powerful promoter of a ubiquitous nature, or which is selective for one type of tissue. The transgenic animals may also be obtained by homologous recombination on embryonic stem cells, transfer of these stem cells to embryos, selection of the chimeras affected at the level of the reproductive lines, and growth of the said chimeras.

The transformed cells as well as the transgenic animals according to the invention can be used in methods of preparing the recombinant polypeptide.

It is now possible to produce recombinant polypeptides in a relatively large quantity by genetic engineering using the cells transformed with expression vectors according to the invention or using transgenic animals according to the invention.

The methods of preparing a polypeptide of the invention in recombinant form, 5 characterized in that they use a vector and/or a cell transformed with a vector according to the invention and/or a transgenic animal comprising one of the said transformed cells according to the invention, are themselves included in the present invention.

Among the said methods of preparing a polypeptide of the invention in recombinant form, the methods of preparation using a vector, and/or a cell transformed with the said vector and/or a transgenic animal comprising one of the said transformed cells, containing a nucleotide sequence encoding a polypeptide of the cellular envelope of *Chlamydia-pneumoniae* or one of its representative fragments, more preferably encoding a polypeptide of the outer cellular envelope of *Chlamydia pneumoniae* or one of its fragment, are preferred.

Among the said methods of preparing a polypeptide of the invention in recombinant form, the methods of preparation using a vector, and/or a cell transformed with the said vector and/or a transgenic animal comprising one of the said transformed cells, containing a nucleotide sequence encoding a Chlamydia pneumoniae secreted polypeptide or one of its representative fragments or encoding a transport polypeptide, a surface exposed polypeptide, a lipoprotein or one of its representative fragments, a polypeptide involved in lipopolysaccharide biosynthesis, a Type III or other secreted polypeptide, a polypeptide containing RGD attachment sites, a cell wall anchored surface polypeptide, a polypeptide not found in Chlamydia trachomatis, a ribosomal polypeptide or a polypeptide involved in secretion, transcription, translation, maturation of proteins, a polypeptide involved in the synthesis of the wall, a polypeptide involved in the virulence, a polypeptide involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, a polypeptide involved in the metabolism of nucleotides, of amino acids, of nucleic acids or of fatty acids of Chlamydia pneumoniae or one of their representative fragments, or a polypeptide specific to Chlamydia pneumoniae, are also preferred.

The recombinant polypeptides obtained as indicated above may be provided either in glycosylated or non-glycosylated form and may or may not have the natural tertiary structure.

A preferred variant consists in producing a recombinant polypeptide fused to a "carrier" protein (chimeric protein). The advantage of this system is that it allows a stabilization and a reduction in proteolysis of the recombinant product, an increase in solubility during renaturation in vitro and/or a simplification of purification when the fusion partner has affinity for a specific ligand.

More particularly, the invention relates to a method of preparing a polypeptide of the invention comprising the following steps:

a) culture of the transformed cells under conditions allowing the expression of a recombinant polypeptide having a nucleic acid sequence according to the invention;

b) where appropriate, recovery of the said recombinant polypeptide.

When the method of preparing a polypeptide of the invention uses a transgenic animal according to the invention, the recombinant polypeptide is then extracted from the said animal.

The subject of the invention is also a polypeptide capable of being obtained by a method 5 of the invention as described above.

The invention also comprises a method of preparing a synthetic polypeptide, characterized in that it uses an amino acid sequence of polypeptides according to the invention.

The invention also relates to a synthetic polypeptide obtained by a method according to the invention.

Polypeptides according to the invention may also be prepared by conventional techniques in the field of peptide synthesis under conditions suitable to produce the polypeptides encoded by the polynucleotide of the invention. This synthesis may be carried out in and recovered from a homogeneous solution or on a solid phase.

For example, the synthesis technique in a homogeneous solution described by 15 Houbenweyl in 1974 may be used.

This method of synthesis consists in successively condensing, in pairs, the successive amino acids in the required order, or in condensing amino acids and fragments previously formed and already containing several amino acids in the appropriate order, or alternatively several fragments thus previously prepared, it being understood that care will have been taken to protect beforehand all the reactive functional groups carried by these amino acids or fragments, with the exception of the amine functional groups of one and the carboxyl functional groups of the other or vice versa, which should normally take part in the formation of the peptide bonds, in particular after activation of the carboxyl functional group, according to methods well known in peptide synthesis.

According to another preferred technique of the invention, the one described by 25 Merrifield is used.

To manufacture a peptide chain according to the Merrifield method, a highly porous polymer resin is used, onto which the first C-terminal amino acid of the chain is attached. This amino acid is attached onto a resin via its carboxyl group and its amine functional group is protected. The amino acids which will constitute the peptide chain are thus attached, one after another, onto the amine group, each time deprotected beforehand, of the portion of the peptide chain already formed, and which is attached to the resin. When the entire peptide chain desired is formed, the protecting groups are removed from the various amino acids constituting the peptide chain and the peptide is detached from the resin with the aid of an acid.

The invention relates, in addition, to hybrid (fusion) polypeptides having at least one polypeptide or one of its representative fragments according to the invention, and a sequence of a polypeptide capable of eliciting an immune response in humans or animals.

Advantageously, the antigenic determinant is such that it is capable of eliciting a humoral

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An antigenic determinant may be identified by screening and/or cellular response. expression libraries of the Chlamydia pneumoniae genome with antibodies contained in the serum of patients infected with a bacterium belonging to the species Chlamydia pneumoniae. An antigenic determinant may comprise a polypeptide or one of its representative fragments according to the 5 invention, in glycosylated form, used in order to obtain immunogenic compositions capable of inducing the synthesis of antibodies directed against multiple epitopes. The said polypeptides or their glycosylated fragments also form part of the invention.

These hybrid molecules may consist, in part, of a carrier molecule for polypeptides or for their representative fragments according to the invention, combined with a portion which may be 10 immunogenic, in particular an epitope of the diphtheria toxin, the tetanus toxin, a hepatitis B virus surface antigen (patent FR 79 21811), the poliomyelitis virus VP1 antigen or any other viral or bacterial toxin or antigen.

The methods of synthesizing the hybrid molecules include the methods used in genetic engineering to construct hybrid nucleotide sequences encoding the desired polypeptide sequences. 15 Reference may be advantageously made, for example, to the technique for producing genes encoding fusion proteins described by Minton in 1984.

The said hybrid nucleotide sequences encoding a hybrid polypeptide as well as the hybrid polypeptides according to the invention, characterized in that they are recombinant polypeptides obtained by the expression of the said hybrid nucleotide sequences, also form part of the invention.

The invention also comprises the vectors characterized in that they contain one of the said hybrid nucleotide sequences. The host cells transformed by the said vectors, the transgenic animals comprising one of the said transformed cells as well as the methods of preparing recombinant polypeptides using the said vectors, the said transformed cells and/or the said transgenic animals of course also form part of the invention.

The polypeptides according to the invention, the antibodies according to the invention described below and the nucleotide sequences according to the invention may advantageously be used in in vitro and/or in vivo methods for the detection and/or the identification of bacteria belonging to the species Chlamydia pneumoniae, in a biological sample (biological tissue or fluid) which is likely to contain them. These methods, depending on the specificity of the polypeptides, of the antibodies 30 and of the nucleotide sequences according to the invention which will be used, may in particular detect and/or identify the bacterial variants belonging to the species Chlamydia pneumoniae as well as the associated microorganisms capable of being detected by the polypeptides, the antibodies and the nucleotide sequences according to the invention which will be chosen. It may, for example, be advantageous to choose a polypeptide, an antibody or a nucleotide sequence according to the 35 invention, which is capable of detecting any bacterium of the Chlamydia family by choosing a polypeptide, an antibody and/or a nucleotide sequence according to the invention which is specific to the family or, on the contrary, it will be most particularly advantageous to target a variant of the species *Chlamydia pneumoniae*, which is responsible, for example, for the induction or the worsening of pathologies specific to the targeted variant, by choosing a polypeptide, an antibody and/or a nucleotide sequence according to the invention which is specific to the said variant.

The polypeptides according to the invention may advantageously be used in a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism, in a biological sample (biological tissue or fluid) which is likely to contain them, characterized in that it comprises the following steps:

- a) bringing this biological sample into contact with a polypeptide or one of its representative fragments according to the invention (under conditions allowing an immunological reaction between
   the said polypeptide and the antibodies which may be present in the biological sample);
  - b) detecting the antigen-antibody complexes which may be formed.

Preferably, the biological sample consists of a fluid, for example a human or animal serum, blood or biopsies.

Any conventional procedure may be used to carry out such a detection of the antigenantibody complexes which may be formed.

By way of example, a preferred method uses immunoenzymatic procedures based on the ELISA technique, immunofluorescence procedures or radioimmunological procedures (RIA), and the like.

Accordingly, the invention also relates to the polypeptides according to the invention, 20 labelled with the aid of a suitable label such as a label of the enzymatic, fluorescent or radioactive type.

Such methods comprise, for example, the following steps:

- deposition of defined quantities of a polypeptide composition according to the invention into the wells of a microtitre plate,
- 25 introduction, into the said wells, of increasing dilutions of serum, or of a different biological sample as defined above, which has to be analysed,
  - incubation of the microplate,

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- introduction, into the wells of the microtitre plate, of labelled antibodies directed against human or animal immunoglobulins, these antibodies having been labelled with the aid of an enzyme selected from those which are capable of hydrolyzing a substrate, thereby modifying the absorption of the radiation of the latter, at least at a defined wavelength, for example at 550 nm,
- detection, by comparison with a control, of the quantity of substrate hydrolyzed.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism, characterized in that it comprises the following components:

- a polypeptide according to the invention,

- where appropriate, the reagents for constituting the medium appropriate for the immunological or specific reaction,
- the reagents allowing the detection of the antigen-antibody complexes produced by the immuno-logical reaction between the polypeptide(s) of the invention and the antibodies which may be present in the biological sample, it being possible for these reagents also to carry a label, or to be capable of being recognized in turn by a labelled reagent, more particularly in the case where the polypeptide according to the invention is not labelled,
- where appropriate, a reference biological sample (negative control) free of antibodies recognized by a polypeptide according to the invention,
- where appropriate, a reference biological sample (positive control) containing a predetermined quantity of antibodies recognized by a polypeptide according to the invention.

According to the invention, the polypeptides, peptides, fusion proteins or other derivatives, or analogs thereof encoded by a polynucleotide sequence in SEQ ID No. 1, may be used as an immunogen to generate antibodies which immunospecifically bind such an immunogen. Such antibodies may include, but are not limited to, polyclonal and monoclonal antibodies, humanized or chimeric antibodies, single chain antibodies, Fab fragments, F(ab')<sub>2</sub> fragments, fragments produced by a Fab expression library, anti-idiotypic (anti-Id) antibodies, and epitope-binding fragments of any of the above. In a specific embodiment, the antibody to a polypeptide, peptide or other derivative, or analog thereof encoded by a polynucleotide sequence in SEQ ID No. 1 is a bispecific antibody (see generally, e.g. Fanger and Drakeman, 1995, Drug News and Perspectives 8: 133-137). Such a bispecific antibody is genetically engineered to recognize both (1) an epitope and (2) one of a variety of "trigger" molecules, e.g. Fc receptors on myeloid cells, and CD3 and CD2 on T cells, that have been identified as being able to cause a cytotoxic T-cell to destroy a particular target. Such bispecific antibodies can be prepared either by chemical conjugation, hybridoma, or recombinant molecular biology techniques known to the skilled artisan.

Various procedures known in the art may be used for the production of polyclonal antibodies to a polypeptide, peptide or other derivative, or analog thereof encoded by a polynucleotide sequence in SEQ ID No. 1. For the production of antibody, various host animals can be immunized by injection with a polypeptide, or peptide or other derivative, or analog thereof, including but not limited to rabbits, mice, rats, etc. Various adjuvants, depending on the host species, may be used to increase the immunological response, including but not limited to Stimulon<sup>TM</sup> QS-21 (Aquila Biopharmaceuticals, Inc., Framingham, MA), MPL<sup>TM</sup> (3-O-deacylated monophosphoryl lipid A; RIBI ImmunoChem Research, Inc., Hamilton, MT), aluminum phosphate, IL-12 (Genetics Institute, Cambridge, MA), Freund's (complete and incomplete), mineral gels such as aluminum hydroxide, surface active substances such as lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, keyhole limpet hemocyanins, dinitrophenol, BCG (bacille Calmette-Guerin), and corynebacterium parvum. Alternatively, polyclonal antibodies may be prepared by purifying, on an affinity column

onto which a polypeptide according to the invention has been previously attached, the antibodies contained in the serum of patients infected with a bacterium belonging to the species Chlamydia pneumoniae.

For preparation of monoclonal antibodies directed toward a polypeptide, peptide or other 5 derivative, or analog, any technique which provides for the production of antibody molecules by continuous cell lines in culture may be used. For example, the hybridoma technique originally developed by Kohler and Milstein (1975, Nature 256:495-497), as well as the trioma technique, the human B-cell hybridoma technique (Kozbor et al., 1983, Immunology Today 4:72), and the EBVhybridoma technique to produce human monoclonal antibodies (Cole et al., 1985, in Monoclonal 10 Antibodies and Cancer Therapy, Alan R. Liss, Inc., pp. 77-96). In an additional embodiment of the invention, monoclonal antibodies can be produced in germ-free animals utilizing technology described in PCT/US90/02545. In another embodiment of the invention, transgenic non-human animals can be used for the production of human antibodies utilizing technology described in WO 98/24893 and WO 96/33735. According to the invention, human antibodies may be used and can be obtained by using 15 human hybridomas (Cote et al., 1983, Proc. Natl. Acad. Sci. U.S.A. 80:2026-2030) or by transforming human B cells with EBV virus in vitro (Cole et al., 1985, in Monoclonal Antibodies and Cancer Therapy, Alan R. Liss, pp. 77-96). In fact, according to the invention, techniques developed for the production of "chimeric antibodies" (Morrison et al., 1984, PROC. NATL. ACAD. SCI. U.S.A. 81:6851-6855; Neuberger et al., 1984, Nature 312:604-608; Takeda et al., 1985, Nature 314:452-454) 20 by splicing the genes from a mouse antibody molecule specific for a polypeptide, peptide or other derivative, or analog together with genes from a human antibody molecule of appropriate biological activity can be used; such antibodies are within the scope of this invention.

According to the invention, techniques described for the production of single chain antibodies (U.S. Patent 4,946,778) can be adapted to produce polypeptide or peptide-specific single chain antibodies. An additional embodiment of the invention utilizes the techniques described for the construction of Fab expression libraries (Huse et al., 1989, Science 246:1275-1281) to allow rapid and easy identification of monoclonal Fab fragments with the desired specificity for polypeptides, derivatives, or analogs.

Antibody fragments which contain the idiotype of the molecule can be generated by 30 known techniques. For example, such fragments include but are not limited to: the F(ab')<sub>2</sub> fragment which can be produced by pepsin digestion of the antibody molecule; the Fab' fragments which can be generated by reducing the disulfide bridges of the F(ab')<sub>2</sub> fragment, the Fab fragments which can be generated by treating the antibody molecule with papain and a reducing agent, and Fv fragments.

In addition, techniques have been developed for the production of chimerized (See, e.g., 35 Boss, M. et al., U.S. Patent No. 4,816,397; and Cabilly, S. et al., U.S. Patent No. 5,585,089 each of which is incorporated herein by reference in its entirety) humanized antibodies (See, e.g., Queen, U.S. Patent No. 5,585,089, which is incorporated herein by reference in its entirety.) An immunoglobulin

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light or heavy chain variable region consists of a "framework" region interrupted by three hypervariable regions, referred to as complementarily determining regions (CDRs). The extent of the framework region and CDRs have been precisely defined (See, "Sequences of Proteins of Immunological Interest", Kabat, E. et al., U.S. Department of Health and Human Services (1983).

Briefly, humanized antibodies are antibody molecules from non-human species having one or more CDRs from the non-human species and a framework from a human immunoglobulin molecule.

The antibodies of the invention may also be labelled in the same manner as described above for the nucleic probes of the invention such as an enzymatic, fluorescent or radioactive type labelling.

- The invention relates, in addition, to a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism in a biological sample, characterized in that it comprises the following steps:
  - a) bringing the biological sample (biological tissue or fluid) into contact with a mono- or polyclonal antibody according to the invention (under conditions allowing an immunological reaction between the said antibodies and the polypeptides of the bacterium belonging to the species Chlamydia pneumoniae or to an associated microorganism which may be present in the biological sample, that is, under conditions suitable for the formation of immune complexes);
  - b) detecting the antigen-antibody complex which may be formed.
- Also falling within the scope of the invention is a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism, characterized in that it comprises the following components:
  - a polyclonal or monoclonal antibody according to the invention, labeled where appropriate;
  - where appropriate, a reagent for constituting the medium appropriate for carrying out the immunological reaction;
  - a reagent allowing the detection of the antigen-antibody complexes produced by the immunological reaction, it being possible for this reagent also to carry a label, or to be capable of being recognized in turn by a labelled reagent, more particularly in the case where the said monoclonal or polyclonal antibody is not labelled;
- 30 where appropriate, reagents for carrying out the lysis of the cells in the sample tested.

The principle of the DNA chip which was explained above may also be used to produce protein "chips" on which the support has been coated with a polypeptide or an antibody according to the invention, or arrays thereof, in place of the DNA. These protein "chips" make it possible, for example, to analyze the biomolecular interactions (BIA) induced by the affinity capture of target analytes onto a support coated, for example, with proteins, by surface plasma resonance (SPR). Reference may be made, for example, to the techniques for coupling proteins onto a solid support which are described in EP 524 800 or to the methods describing the use of biosensor-type protein

chips such as the BIAcore-type technique (Pharmacia) (Arlinghaus et al., 1997, Krone et al., 1997, Chatelier et al., 1995). These polypeptides or antibodies according to the invention, capable of specifically binding antibodies or polypeptides derived from the sample to be analysed, may thus be used in protein chips for the detection and/or the identification of proteins in samples. The said protein chips may in particular be used for infectious diagnosis and may preferably contain, per chip, several polypeptides and/or antibodies of the invention of different specificity, and/or polypeptides and/or antibodies capable of recognizing microorganisms different from *Chlamydia pneumoniae*.

Accordingly, the subject of the present invention is also the polypeptides and the antibodies according to the invention, characterized in that they are immobilized on a support, in particular of a protein chip.

The protein chips, characterized in that they contain at least one polypeptide or one antibody according to the invention immobilized on the support of the said chip, also form part of the invention.

The invention comprises, in addition, a protein chip according to the invention, 15 characterized in that it contains, in addition, at least one polypeptide of a microorganism different from *Chlamydia pneumoniae* or at least one antibody directed against a compound of a microorganism different from *Chlamydia pneumoniae*, immobilized on the support of the said chip.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism, or for the detection and/or the identification of a microorganism characterized in that it comprises a protein chip according to the invention.

The subject of the present invention is also a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism in a biological sample, characterized in that it uses a nucleotide sequence according to the invention.

More particularly, the invention relates to a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism in a biological sample, characterized in that it comprises the following steps:

- a) where appropriate, isolation of the DNA from the biological sample to be analysed, or optionally production of a cDNA from the RNA in the biological sample;
- specific amplification of the DNA of bacteria belonging to the species Chlamydia pneumoniae or to an associated microorganism with the aid of at least one primer according to the invention;
- c) detection of the amplification products.

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These may be detected, for example, by the molecular hybridization technique using a nucleic probe according to the invention. This probe will be advantageously labelled with a nonradioactive (cold probe) or radioactive element.

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For the purposes of the present invention, "DNA in the biological sample" or "DNA contained in the biological sample" will be understood to mean either the DNA present in the biological sample considered, or optionally the cDNA obtained after the action of a reverse transcriptase-type enzyme on the RNA present in the said biological sample.

- Another aim of the present invention consists in a method according to the invention, characterized in that it comprises the following steps:
  - a) bringing a nucleotide probe according to the invention into contact with a biological sample, the DNA contained in the biological sample having, where appropriate, been previously made accessible to hybridization, under conditions allowing the hybridization of the probe to complementary base pairs of the DNA of a bacterium belonging to the species Chlamydia pneumoniae or to an associated-microorganism;
  - b) detecting the hybridization complex formed between the nucleotide probe and the DNA in the biological sample.

The present invention also relates to a method according to the invention, characterized in that it comprises the following steps:

- bringing a nucleotide probe immobilized on a support according to the invention into contact with a biological sample, the DNA in the sample having, where appropriate, been previously made accessible to hybridization, under conditions allowing the hybridization of the probe to the DNA of a bacterium belonging to the species *Chlamydia pneumoniae* or to an associated microorganism;
- b) bringing the hybrid formed between the nucleotide probe immobilized on a support and the DNA contained in the biological sample, where appropriate after removal of the DNA in the biological sample which has not hybridized with the probe, into contact with a labelled nucleotide probe according to the invention;
- 25 c) detecting the new hybrid formed in step b).

According to an advantageous embodiment of the method for the detection and/or the identification defined above, it is characterized in that, prior to step a), the DNA in the biological sample is primer-extended and/or amplified beforehand with the aid of at least one primer according to the invention.

- The invention relates, in addition, to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia pneumoniae* or to an associated microorganism, characterized in that it comprises the following components:
  - a) a nucleotide probe according to the invention;
  - b) where appropriate, the reagents necessary for carrying out a hybridization reaction;
- where appropriate, at least one primer according to the invention as well as the reagents (e.g., polymerase and/or deoxynucleotide triphosphates) necessary for a DNA amplification reaction.

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The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species Chlamydia pneumoniae or to an associated microorganism, characterized in that it comprises the following components:

- a nucleotide probe, called capture probe, according to the invention;
- an oligonucleotide probe, called detection probe, according to the invention; 5 b)
  - where appropriate, at least one primer according to the invention as well as the reagents (e.g., polymerase and/or deoxynucleotide triphosphates) necessary for a DNA amplification reaction.

The invention also relates to a kit or set for the detection and/or the identification of 10 bacteria belonging to the species Chlamydia pneumoniae or to an associated microorganism, characterized in that it comprises the following components:

- at least one primer according to the invention;
- where appropriate, the reagents necessary for carrying out a DNA amplification reaction; b)
- where appropriate, a component which makes it possible to check the sequence of the amplified c) fragment, more particularly an oligonucleotide probe according to the invention.

The invention relates, in addition, to a kit or set for the detection and/or the identification of bacteria belonging to the species Chlamydia pneumoniae or to an associated microorganism, or for the detection and/or the identification of a microorganism characterized in that it comprises a DNA chip according to the invention.

The invention also relates to a method or to a kit or set according to the invention for the detection and/or the identification of bacteria belonging to the species Chlamydia pneumoniae, characterized in that the said primer and/or the said probe according to the invention are chosen from the nucleotide sequences specific to the species Chlamydia pneumoniae, in that the said polypeptides according to the invention are chosen from the polypeptides specific to the species Chlamydia 25 pneumoniae and in that the said antibodies according to the invention are chosen from the antibodies directed against the polypeptides according to the invention chosen from the polypeptides specific to the species Chlamydia pneumoniae.

Preferably, the said method or the said kit or set above according to the invention, for the detection and/or the identification of bacteria belonging to the species Chlamydia pneumoniae is 30 characterized in that the said primer and/or the said probe or the said polypeptides are chosen from the nucleotide sequences or polypeptides according to the invention which have been identified as being specific to the species Chlamydia pneumoniae and in that the said antibodies according to the invention are chosen from the antibodies directed against the polypeptides according to the invention chosen from the polypeptides identified as being specific to the species Chlamydia pneumoniae.

The invention relates, in addition, to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of a condition caused by, cardiovascular diseases, preferably linked to the presence of atheroma, which are induced or worsened by a Chlamydia pneumoniae infection.

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The invention also relates to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of conditions caused by, respiratory diseases induced or worsened by a Chlamydia pneumoniae infection; preferably, the said respiratory disease is asthma.

According to another aspect, the subject of the invention is the use of polypeptides according to the invention, of cells transformed with a vector according to the invention and/or of transformed animals according to the invention, for the biosynthesis or the biodegradation of organic or inorganic compounds.

As has been mentioned above, the nucleotide sequences of the invention were identified 10 by homology with sequences known to encode, for example, polypeptides or fragments of enzymatic polypeptides involved in the biosynthesis or the biodegradation of organic or inorganic molecules.

It is thus possible to use the said polypeptides of the invention in a similar manner for the biosynthesis or the biodegradation of organic or inorganic compounds of industrial or therapeutic interest (called compounds of interest).

Among these polypeptides, there may be mentioned in particular the enzymes involved in metabolism, such as the proteolytic enzymes, amino transferases, glucose metabolism, or the enzymes which may be used in the biosynthesis of sugars, amino acids, fatty acids, polypeptides, nucleotides, nucleic acids or any other organic or inorganic compound or in the biodegradation of organic or inorganic compounds.

Among these polypeptides, there may be mentioned, in addition, the mutated or modified enzymes corresponding to mutated or modified polypeptides according to the invention which may also be used for the biosynthesis or the biodegradation of organic or inorganic compounds at the industrial level, such as, for example, the production of compounds of interest, the reprocessing of manufacturing residues applied to the food industries, to the papermaking industry or to the chemical 25 and pharmaceutical industries.

The methods of biosynthesis or biodegradation of organic or inorganic compounds, characterized in that they use a polypeptide or one of its representative fragments according to the invention, transformed cells according to the invention and/or a transformed animal according to the invention, also form part of the invention.

The invention relates, in addition, to the use of a nucleotide sequence according to the invention, of a polypeptide according to the invention, of an antibody according to the invention, of a cell according to the invention, and/or of a transformed animal according to the invention, for the selection of an organic or inorganic compound capable of modulating, regulating, inducing or inhibiting the expression of genes, and/or of modifying the cellular replication of eukaryotic or 35 prokaryotic cells or capable of inducing, inhibiting or worsening the pathologies linked to an infection by Chlamydia pneumoniae or one of its associated microorganisms.

The invention also comprises screening assays that comprise methods of selecting

compounds capable of binding to a polypeptide, fusion polypeptide or one of its representative fragments according to the invention, capable of binding to a nucleotide sequence according to the invention, or capable of recognizing an antibody according to the invention, and/or capable of modulating, regulating, inducing or inhibiting the expression of genes, and/or of modifying the growth or the cellular replication of eukaryotic or prokaryotic cells, or capable of inducing, inhibiting or worsening, in an animal or human organism, the pathologies linked to an infection by *Chlamydia pneumoniae* or one of its associated microorganisms, characterized in that it comprises the following steps:

- a) bringing the said compound into contact with the said polypeptide, the said nucleotide
   sequence, with a transformed cell according to the invention and/or administering the said compound to a transformed animal according to the invention;
- b) determining the capacity of the said compound to bind with the said polypeptide or the said nucleotide sequence, or to modulate, regulate, induce or inhibit the expression of genes, or to modulate growth or cellular replication, or to induce, inhibit or worsen in the said transformed animal,
   the pathologies linked to an infection by *Chlamydia pneumoniae* or one of its associated microorganisms.

The transformed cells and/or animals according to the invention may advantageously serve as a model and may be used in methods for studying, identifying and/or selecting compounds capable of being responsible for pathologies induced or worsened by *Chlamydia pneumoniae*, or capable of preventing and/or of treating these pathologies such as, for example, cardiovascular or respiratory diseases. In particular, the transformed host cells, in particular bacteria of the *Chlamydia* family whose transformation with a vector according to the invention may, for example, increase or inhibit its infectivity, or modulate the pathologies usually induced or worsened by the infection, may be used to infect animals in which the onset of pathologies will be monitored. These nontransformed animals, infected for example with transformed *Chlamydia* bacteria, may serve as a study model. In the same manner, the transformed animals according to the invention may, for example, exhibit predispositions to cardiovascular and/or respiratory diseases and thus be used in methods for selecting compounds capable of preventing and/or of treating the said diseases. The said methods using the said transformed cells and/or transformed animals form part of the invention.

The compounds capable of being selected may be organic compounds such as polypeptides or carbohydrates or any other organic or inorganic compounds already known, or new organic compounds produced using molecular modeling techniques and obtained by chemical or biochemical synthesis, these techniques being known to persons skilled in the art.

The said selected compounds may be used to modulate the growth and/or the cellular replication of *Chlamydia pneumoniae* or any other associated microorganism and thus to control infection by these microorganisms. The said compounds according to the invention may also be used to modulate the growth and/or the cellular replication of all eukaryotic or prokaryotic cells, in

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infectious microorganisms, for which the said compounds particular cells tumour and will prove active, the methods which make it possible to determine the said modulations being well known to persons skilled in the art.

Compound capable of modulating the growth of a microorganism is understood to 5 designate any compound which makes it possible to act, to modify, to limit and/or to reduce the development, the growth, the rate of proliferation and/or the viability of the said microorganism.

This modulation may be achieved, for example, by an agent capable of binding to a protein and thus of inhibiting or of potentiating its biological activity, or capable of binding to a membrane protein of the outer surface of a microorganism and of blocking the penetration of the said 10 microorganism into the host cell or of promoting the action of the immune system of the infected organism directed against the said microorganism. This modulation may also be achieved by an agent capable of binding to a nucleotide sequence of a DNA or RNA of a microorganism and of blocking, for example, the expression of a polypeptide whose biological or structural activity is necessary for the growth or for the reproduction of the said microorganism.

Associated microorganism is understood to designate in the present invention any microorganism whose gene expression may be modulated, regulated, induced or inhibited, or whose growth or cellular replication may also be modulated by a compound of the invention. Associated microorganism is also understood to designate in the present invention any microorganism containing nucleotide sequences or polypeptides according to the invention. These microorganisms may, in some 20 cases, contain polypeptides or nucleotide sequences identical or homologous to those of the invention may also be detected and/or identified by the detection and/or identification methods or kit according to the invention and may also serve as a target for the compounds of the invention.

The invention relates to the compounds capable of being selected by a method of selection according to the invention.

The invention also relates to a pharmaceutical composition comprising a compound chosen from the following compounds:

a nucleotide sequence according to the invention;

a polypeptide according to the invention;

a vector according to the invention;

30 an antibody according to the invention; and

a compound capable of being selected by a method of selection according to the invention, optionally in combination with a pharmaceutically acceptable vehicle.

An effective quantity is understood to designate a sufficient quantity of the said compound or antibody, or of a polypeptide of the invention, which makes it possible to modulate the 35 growth of Chlamydia pneumoniae or of an associated microorganism.

The invention also relates to a pharmaceutical composition comprising one or more polypeptides according to the invention and/or one or more fusion polypeptides according to the

invention. Such compositions further comprise a pharmaceutically acceptable carrier or vehicle. Pharmaceutical compositions include compositions that comprise a polypeptide or fusion polypeptide that immunoreacts with seropositive serum of an individual infected with Chlamydia pneumoniae. In one embodiment, a pharmaceutical composition according to the invention can be utilized for the 5 prevention or the treatment of an infection by a bacterium belonging to the species Chlamydia pneumoniae or by an associated microorganism.

The invention relates, in addition, to an immunogenic composition or a vaccine composition, characterized in that it comprises one or more polypeptides according to the invention and/or one or more hybrid (fusion) polypeptides according to the invention. Such compositions 10 further comprise a pharmaceutically acceptable carrier or vehicle. Immunogenic compositions or fusion polypeptide include compositions that comprise a polypeptide that immunoreacts with seropositive serum of an individual infected with Chlamydia pneumoniae.

Immunogenic or vaccine compositions can also comprise DNA immunogenic or vaccine compositions comprising polynucleotide sequences of the invention operatively associated with a 15 regulatory sequence that controls gene expression. Such compositions can include compositions that direct expression of a neutralizing epitope of Chlamydia pneumoniae.

The invention also comprises the use of a transformed cell according to the invention, for the preparation of a vaccine composition.

The invention also relates to a vaccine composition, characterized in that it contains a 20 nucleotide sequence according to the invention, a vector according to the invention and/or a transformed cell according to the invention.

The invention also relates to the vaccine compositions according to the invention, for the prevention or the treatment of an infection by a bacterium belonging to the species Chlamydia pneumoniae or by an associated microorganism.

The invention also relates to the use of DNA encoding polypeptides of Chlamydia pneumoniae, in particular antigenic determinants, to be formulated as vaccine compositions. In accordance with this aspect of the invention, the DNA of interest is engineered into an expression vector under the control of regulatory elements, which will promote expression of the DNA, i.e., promoter or enhancer elements. In one preferred embodiment, the promoter element may be cell-30 specific and permit substantial transcription of the DNA only in predetermined cells. The DNA may be introduced directly into the host either as naked DNA (U.S. Patent No. 5,679,647 incorporated herein by reference in their entirety) or formulated in compositions with other agents which may facilitate uptake of the DNA including viral vectors, i.e., adenovirus vectors, or agents which facilitate immunization, such as bupivicaine and other local anesthetics (U.S. Patent 5,593,972 incorporated 35 herein by reference in their entirety), saponins (U.S. Patent 5,739,118 incorporated herein by reference in their entirety) and cationic polyamines (published international application WO 96/10038 incorporated herein by reference in their entirety).

The DNA sequence encoding the antigenic polypeptide and regulatory element may be inserted into a stable cell line or cloned microorganism, using techniques, such as targeted homologous recombination, which are well known to those of skill in the art, and described e.g., in Chappel, U.S. Patent No. 4,215,051; Skoultchi, WO 91/06667 each of which is incorporated herein by reference in its entirety.

Such cell lines and microorganisms may be formulated for vaccine purposes. In yet another embodiment, the DNA sequence encoding the antigenic polypeptide and regulatory element may be delivered to a mammalian host and introduced into the host genome via homologous recombination (See, Chappel, U.S. Patent No. 4,215,051; Skoultchi, WO 91/06667 each of which is incorporated herein by reference in its entirety.

Preferably, the immunogenic and/or vaccine compositions according to the invention intended for the prevention and/or the treatment of an infection by Chlamydia pneumoniae or by an associated microorganism will be chosen from the immunogenic and/or vaccine compositions comprising a polypeptide or one of its representative fragments corresponding to a protein, or one of its representative fragments, of the cellular envelope of Chlamydia pneumoniae. The vaccine compositions comprising nucleotide sequences will also preferably comprise nucleotide sequences encoding a polypeptide or one of its representative fragments corresponding to a protein, or one of its representative fragments, of the cellular envelope of Chlamydia pneumoniae.

Among these preferred immunogenic and/or vaccine compositions, the most preferred are
those comprising a polypeptide or one of its representative fragments, or a nucleotide sequence or one
of its representative fragments whose sequences are chosen from the nucleotide or amino acid
sequences identified in this functional group and listed above.

The polypeptides of the invention or their representative fragments entering into the immunogenic compositions according to the invention may be selected by techniques known to persons skilled in the art, such as for example on the capacity of the said polypeptides to stimulate T cells, which results, for example, in their proliferation or the secretion of interleukins, and which leads to the production of antibodies directed against the said polypeptides.

In mice, in which a weight dose of the vaccine composition comparable to the dose used in humans is administered, the antibody reaction is tested by collecting serum followed by a study of the formation of a complex between the antibodies present in the serum and the antigen of the vaccine composition, according to the customary techniques.

According to the invention, the said vaccine compositions will be preferably in combination with a pharmaceutically acceptable vehicle and, where appropriate, with one or more appropriate immunity adjuvants.

Various types of vaccines are currently available for protecting humans against infectious diseases: attenuated live microorganisms (M. bovis - BCG for tuberculosis), inactivated microorganisms (influenza virus), acellular extracts (Bordetella pertussis for whooping cough),

recombinant proteins (hepatitis B virus surface antigen), polysaccharides (pneumococci). Experiments are underway on vaccines prepared from synthetic peptides or from genetically modified microorganisms expressing heterologous antigens. Even more recently, recombinant plasmid DNAs carrying genes encoding protective antigens were proposed as an alternative vaccine strategy. This type of vaccination is carried out with a particular plasmid derived from an *E. coli* plasmid which does not replicate *in vivo* and which encodes only the vaccinal protein. Animals were immunized by simply injecting the naked plasmid DNA into the muscle. This technique leads to the expression of the vaccine protein *in situ* and to a cell-type (CTL) and a humoral type (antibody) immune response. This double induction of the immune response is one of the main advantages of the technique of vaccination with naked DNA.

The vaccine compositions of the present invention can be evaluated in in vitro and in vivo animal models prior to host, e.g., human, administration. For example, in vitro neutralization assays such as those described by Peterson et al. (1988) can be utilized. The assay described by Peterson et al. (1988) is suitable for testing vaccine compositions directed toward either Chlamydia pneumoniae or Chlamydia trachomatis.

Briefly, hyper-immune antisera is diluted in PBS containing 5% guinea pig serum, as a complement source. Chlamydiae (10<sup>4</sup> IFU; infectious units) are added to the antisera dilutions. The antigen-antibody mixtures are incubated at 37EC for 45 minutes and inoculated into duplicate confluent Hep-2 or HeLa cell monolayers contained in glass vials (e.g., 15 by 45 mm), which have been washed twice with PBS prior to inoculation. The monolayer cells are infected by centrifugation at 1000X g for 1 hour followed by stationary incubation at 37E for 1 hour. Infected monolayers are incubated for 48 or 72 hours, fixed and stained with a Chlamydiae specific antibody, such as anti-MOMP for C.trachomatis, etc. IFUs are counted in ten fields at a magnification of 200X. Neutralization titer is assigned based on the dilution that gives 50% inhibition as compared to control monolayers/IFU.

The efficacy of vaccine compositions can be determined *in vivo* by challenging animal models of *Chlamydia pneumoniae* infection, <u>e.g.</u>, mice or rabbits, with the vaccine compositions. For example, *in vivo* vaccine composition challenge studies can be performed in the murine model of *Chlamydia pneumonia* infection described by Moazed et al. (1997). Briefly, male homozygous apoE deficient and/or C57 BL/6J mice are immunized with vaccine compositions. Post-vaccination, the mice are mildly sedated by subcutaneous injection of a mixture of ketamine and xylazine, and inoculated intranasally with a total volume of 0.03-0.05 ml of organisms suspended in SPG medium or with SPG alone. The inoculations of *Chlamydia pneumoniae* are approximately 3x10<sup>7</sup> IFU/mouse. The mice are inoculated with *Chlamydia pneumoniae* at 8, 10, and 12 weeks of age. Tissues are then collected from the lung, spleen, heart, etc. at 1-20 weeks after the first inoculation. The presence of organisms is scored using PCR, histology and immunocytochemistry, or by quantitative culture/IFU after tissue homogenization.

Alternatively, in vivo vaccine composition challenge studies can be performed in the rabbit model of Chlamydia pneumoniae described by Laitinen et al. (1997). Briefly, New Zealand white rabbits (5 months old) are immunized with the vaccine compositions. Post-vaccination, the rabbits are sedated with Hypnorm, 0.3 ml/Kg of body weight, intramuscularly, and inoculated intranasally with a total of 0.5 ml of Chlamydia pneumoniae suspended in SPG medium or with SPG alone. The inoculations of Chlamydia pneumoniae are approximately 3x10<sup>7</sup> IFU/rabbit. The rabbits are reinfected in the same manner and with the same dose 3 weeks after the primary inoculation. Tissues are then collected 2 weeks after the primary infection and 1, 2, and 4 weeks after the reinfection. The presence of Chlamydia pneumoniae is scored using PCR, histology and immunocytochemistry, or by quantitative culture/IFU after tissue homogenization.

The vaccine compositions comprising nucleotide sequences or vectors into which the said sequences are inserted are in particular described in International Application No. WO 90/11092 and also in International Application No. WO 95/11307.

The nucleotide sequence constituting the vaccine composition according to the invention may be injected into the host after having been coupled to compounds which promote the penetration of this polynucleotide inside the cell or its transport up to the cell nucleus. The resulting conjugates may be encapsulated into polymeric microparticles, as described in International Application No. WO 94/27238 (Medisorb Technologies International).

According to another embodiment of the vaccine composition according to the invention, the nucleotide sequence, preferably a DNA, is complexed with the DEAE-dextran (Pagano et al., 1967) or with nuclear proteins (Kaneda et al., 1989), with lipids (Felgner et al., 1987) or encapsulated into liposomes (Fraley et al., 1980) or alternatively introduced in the form of a gel facilitating its transfection into the cells (Midoux et al., 1993, Pastore et al., 1994). The polynucleotide or the vector according to the invention may also be in suspension in a buffer solution or may be combined with liposomes.

Advantageously, such a vaccine will be prepared in accordance with the technique described by Tacson et al. or Huygen et al. in 1996 or alternatively in accordance with the technique described by Davis et al. in International Application No. WO 95/11307.

Such a vaccine may also be prepared in the form of a composition containing a vector according to the invention, placed under the control of regulatory elements allowing its expression in humans or animals. It is possible, for example, to use, as vector for the *in vivo* expression of the polypeptide antigen of interest, the plasmid pcDNA3 or the plasmid pcDNA1/neo, both marketed by Invitrogen ® & D Systems, Abingdon, United Kingdom). It is also possible to use the plasmid V1Jns.tPA, described by Shiver et al. in 1995. Such a vaccine will advantageously comprise, in addition to the recombinant vector, a saline solution, for example a sodium chloride solution.

The immunogenic compositions of the invention can also be utilized as part of methods for immunization, wherein such methods comprise administering to a host, e.g., a human host, an

immunizing amount of the immunogenic compositions of the invention. In a preferred embodiment, the method of immunizing is a method of immunizing against Chlamydia pneumoniae.

A pharmaceutically acceptable vehicle is understood to designate a compound or a combination of compounds entering into a pharmaceutical or vaccine composition which does not 5 cause side effects and which makes it possible, for example, to facilitate the administration of the active compound, to increase its life and/or its efficacy in the body, to increase its solubility in solution or alternatively to enhance its preservation. These pharmaceutically acceptable vehicles are well known and will be adapted by persons skilled in the art according to the nature and the mode of administration of the active compound chosen.

As regards the vaccine formulations, these may comprise appropriate immunity adjuvants which are known to persons skilled in the art, such as, for example, aluminum hydroxide, a representative of the family of muramyl peptides such as one of the peptide derivatives of N-acetylmuramyl, a bacterial lysate, or alternatively incomplete Freund's adjuvant, Stimulon™ QS-21 (Aquila Biopharmaceuticals, Inc., Framingham, MA), MPL™ (3-O-deacylated monophosphoryl lipid A; RIBI 15 ImmunoChem Research, Inc., Hamilton, MT), aluminum phosphate, IL-12 (Genetics Institute, Cambridge, MA).

Preferably, these compounds will be administered by the systemic route, in particular by the intravenous route, by the intranasal, intramuscular, intradermal or subcutaneous route, or by the oral route. More preferably, the vaccine composition comprising polypeptides according to the 20 invention will be administered several times, spread out over time, by the intradermal or subcutaneous route.

Their optimum modes of administration, dosages and galenic forms may be determined according to criteria which are generally taken into account in establishing a treatment adapted to a patient, such as for example the patient's age or body weight, the seriousness of his general condition, 25 tolerance of the treatment and the side effects observed.

The invention comprises the use of a composition according to the invention for the treatment or the prevention of cardiovascular diseases, preferably linked to the presence of atheroma, which are induced or worsened by Chlamydia pneumoniae.

Finally, the invention comprises the use of a composition according to the invention for 30 the treatment or the prevention of respiratory diseases which are induced or worsened by the presence of Chlamydia pneumoniae, preferably asthma.

Other characteristics and advantages of the invention appear in the following examples and figures:

### 35 Legend to the figures:

Figure 1: Line for the production of Chlamydia pneumoniae sequences

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Figure 2: Analysis of the sequences and assembling

Figure 3: Finishing techniques

Figure 3a): Assembly map

Figure 3b): Determination and use of the orphan ends of the contigs

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### **EXAMPLES**

## Experimental procedures

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### Cells

The Chlamydia pneumoniae strain (CM1) used by the inventors is obtained from ATCC (American Culture Type Collection) where it has the reference number ATCC 1360-VR.

It is cultured on HeLa 229 cells, obtained from the American Type Culture Collection, under the reference ATCC CCL-2.1.

### Culture of the cells

The HeLa ATCC CCL-2.1 cells are cultured in 75-ml cell culture flasks (Corning). The culture medium is Dulbecco's modified cell culture medium (Gibco BRL No. 04101965) supplemented with MEM amino acids (Gibco BRL - No. 04301140) L (5 ml per 500 ml of medium) and 5% foetal calf serum (Gibco BRL No. 10270 batch 40G8260K) without antibiotics or antifungals.

The cell culture stock is maintained in the following manner. The cell cultures are examined under an inverted microscope. 24 hours after confluence, each cellular lawn is washed with PBS (Gibco BRL No. 04114190), rinsed and then placed for 5 min in an oven in the presence of 3 ml of trypsine (Gibco BRL No. 25200056). The cellular lawn is then detached and then resuspended in 120 ml of culture medium, the whole is stirred in order to make the cellular suspension homogeneous. 30 ml of this suspension are then distributed per cell culture flask. The flasks are kept in a CO<sub>2</sub> oven (5%) for 48 hours at a temperature of 37°C. The cell stock is maintained so as to have available daily 16 flasks of subconfluent cells. It is these subconfluent cells which will be used so as to be infected with Chlamydia. 25-ml cell culture flasks are also used, these flasks are prepared in a similar manner but the volumes used for maintaining the cells are the following: 1 ml of trypsine, 28 ml of culture medium to resuspend the cells, 7 ml of culture medium are used per 25-ml flask.

### Infection of the cells with Chlamydia

Initially, the Chlamydiae are obtained frozen from ATCC (-70°C), in suspension in a volume of 1 ml. This preparation is slowly thawed, 500 µl are collected and brought into contact with subconfluent cells, which are obtained as indicated above, in a 25-ml cell culture flask, containing 1 ml of medium, so as to cover the cells. The flask is then centrifuged at 2000 rpm in a "swing" rotor for microtitre plates, the centrifuge being maintained at a temperature of 35°C. After centrifugation,

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the two flasks are placed in an oven at 35°C for three hours. 6 ml of culture medium containing cycloheximide (1  $\mu$ g/ml) are then added and the flask is stored at 35°C. After 72 hours, the level of infection is evaluated by direct immunofluorescence and by the cytopathogenic effect caused to the cells.

### Direct immunofluorescence

Starting with infected cells, which were obtained as indicated above, a cellular smear is deposited with a Pasteur pipette on a microscope slide. The cellular smear is fixed with acetone for 10 minutes; after draining the acetone, the smear is covered with 30 µl of murine monoclonal antibodies directed against MOMP (major outer membrane protein) of Chlamydia (Syva, Biomérieux) labelled with fluorescein isothiocyanate. The whole is then incubated in a humid chamber at a temperature of 37°C. The slides are then rinsed with water, slightly dried, and then after depositing a drop of mounting medium, a coverslip is mounted before reading. The reading is carried out with the aid of a fluorescence microscope equipped with the required filters (excitation at 490 nm, emission at 520 nm).

### Harvesting of the Chlamydia pneumoniae

After checking the infection by direct immunofluorescence, carried out as indicated above, the culture flasks are opened under a sterile cabinet, sterile glass beads with a diameter of the order of a millimeter are placed in the flask. The flask is closed and then vigorously stirred while being maintained horizontally, the cellular lawn at the bottom, so that the glass beads can have a mechanical action on the cellular lawn. Most of the cells are thus detached or broken; the effect of the stirring is observed under an optical microscope so as to ensure proper release of Chlamydiae.

# Large-scale infection of the cell cultures

The product of the Chlamydiae harvest (culture medium and cellular debris) is collected with a pipette, and distributed into three cell culture flasks containing subconfluent HeLa ATCC CCL-2.1 cells, obtained as indicated above. The cells thus inoculated are placed under gentle stirring (swing) in an oven at 35°C. After one hour, the flasks are kept horizontally in an oven so that the culture medium covers the cells for 3 hours. 30 ml of culture medium containing actydione (1 μg/ml) are then added to each of the flasks. The culture flasks are then stored at 35°C for 72 hours. The cells thus infected are examined under an optical microscope after 24 hours, the cytopathogenic effect is evaluated by the appearance of cytoplasmic inclusions which are visible under an inverted optical microscope. After 72 hours, the vacuoles containing the Chlamydiae occupy the cytoplasm of the cell and push the cell nucleus sideways. At this stage, numerous cells are spontaneously destroyed and have left free elementary bodies in the culture medium. The Chlamydiae are harvested as described above and are either frozen at -80°C or used for another propagation.

### Purification of the Chlamydiae

The product of the Chlamydia harvests is stored at -80°C and thawed on a water bath at

room temperature. After thawing, each tube is vigorously stirred for one minute and immersed for one minute in an ultrasound tank (BRANSON 1200); the tubes are then stirred by inverting before being centrifuged for 5 min at 2000 rpm. The supernatant is carefully removed and kept at cold temperature (ice). The supernatant is vigorously stirred and then filtered on nylon filters having pores 5 of 5 microns in diameter on a support (Nalgene) allowing a delicate vacuum to be established under the nylon filter. For each filtration, three nylon filters are superposed; these filters are replaced after every 40 ml of filtrate. Two hundred milliliters of filtration product are kept at cold temperature, and then after stirring by inverting, are centrifuged at 10,000 rpm for 90 min, the supernatant is removed and the pellet is taken up in 10 ml of 10 mM Tris, vigorously vortexed and then centrifuged at 10 10,000 rpm for 90 min. The supernatant is removed and the pellet is taken up in a buffer (20 mM Tris pH 8.0, 50 mM KCl, 5 mM MgCl<sub>2</sub>) to which 800 units of DNAse I (Boehringer) are added. The whole is kept at 37°C for one hour. One ml of 0.5 M EDTA is then added, the whole is vortexed and frozen at -20°C.

### Preparation of the DNA

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The Chlamydiae purified above are thawed and subjected to a proteinase K (Boehringer) digestion in a final volume of 10 ml. The digestion conditions are the following: 0.1 mg/ml proteinase K, 0.1 x SDS at 55EC, stirring every 10 min. The product of digestion is then subjected to a double extraction with phenol-chloroform, two volumes of ethanol are added and the DNA is directly recovered with a Pasteur pipette having one end in the form of a hook. The DNA is dried on the edge 20 of the tube and then resuspended in 500 μl of 2 mM Tris pH 7.5. The DNA is stored at 4°C for at least 24 hours before being used for the cloning.

### Cloning of the DNA

After precipitation, the DNA is quantified by measuring the optical density at 260 nm. Thirty µg of Chlamydia DNA are distributed into 10 tubes of 1.5 ml and diluted in 300 µl of water. 25 Each of the tubes is subjected to 10 applications of ultrasound lasting for 0.5 sec in a sonicator (unisonix XL2020). The contents of the 10 tubes are then grouped and concentrated by successive extractions with butanol (Sigma B1888) in the following manner: two volumes of butanol are added to the dilute DNA mixture. After stirring, the whole is centrifuged for five minutes at 2500 rpm and the butanol is removed. This operation is repeated until the volume of the aqueous phase is less than 1 ml. 30 The DNA is then precipitated in the presence of ethanol and of 0.5 M sodium acetate pH 5.4, and then centrifuged for thirty minutes at 15,000 rpm at cold temperature (4°C). The pellet is washed with 75% ethanol, centrifuged for five minutes at 15,000 rpm and dried at room temperature. A tenth of the preparation is analysed on a 0.8% agarose gel. Typically, the size of the DNA fragments thus prepared is between 200 and 8000 base pairs.

To allow the cloning of the DNA obtained, the ends are repaired. The DNA is distributed in an amount of 10 µg/tube, in the following reaction medium: 100 µl final volume, 1 × buffer WO 99/27105 PCT/IB98/01890

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(Biolabs 201L), 0.5 μl BSA 0.05 mg/ml, 0.1 mM dATP, 0.1 mM each of dGTP, dCTP or dTTP, 60,000 IU T4 DNA polymerase. The reaction is incubated for thirty minutes at 16°C. The contents of each of the tubes are then grouped before carrying out an extraction with phenol-chloroform and then precipitating the aqueous phase as described above. After this step, the DNA thus prepared is phosphorylated. For that, the DNA is distributed into tubes in an amount of 10 μg per tube, and then in a final volume of 50 μl, the reaction is prepared in the following manner: 1 mM ATP, 1 × kinase buffer, 10 IU T4 polynucleotide kinase (Biolabs 201L). The preparation is incubated for thirty minutes at 37°C. The contents of the tubes are combined and a phenol-chloroform extraction and then a precipitation are carried out in order to precipitate the DNA. The latter is then suspended in 1 μl of water and then the DNA fragments are separated according to their size on a 0.8% agarose gel (1 × TAE). The DNA is subjected to an electric field of 5 V/cm and then visualized on a UV table. The fragments whose size varies between 1200 and 2000 base pairs are selected by cutting out the gel. The gel fragment thus isolated is placed in a tube and then the DNA is purified with the Qiaex kit (20021 Qiagen), according to the procedure provided by the manufacturer.

# Preparation of the vector

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14 μg of the cloning vector pGEM-5Zf (Proméga P2241) are diluted in a final volume of 150 μl and are subjected to digestion with the restriction enzyme EcoRV 300 IU (Biolabs 195S) according to the protocol and with the reagents provided by the manufacturer. The whole is placed at 37°C for 150 min and then distributed in the wells of a 0.8% agarose gel subjected to an electric field of 5 V/cm. The linearized vector is visualized on a UV table, isolated by cutting out the gel and then purified by the Qiaex kit (Qiagen 20021) according to the manufacturer's recommendations. The purification products are grouped in a tube, the volume is measured and then half the volume of phenol is added and the whole is vigorously stirred for 1 min. Half the volume of chloroform-isoamyl alcohol 24:1 is added and vigorously stirred for 1 min. The whole is centrifuged at 15,000 rpm for 5 min at 4°C, the aqueous phase is recovered and transferred into a tube. The DNA is precipitated in the presence of 0.3 M sodium acetate, pH 5.4 and 3 volumes of ethanol and placed at -20°C for 1 hour. The DNA is then centrifuged at 15,000 rpm for 30 min at 4°C, the supernatant is removed while preserving the pellet, washed twice with 70% ethanol. After drying at room temperature, the DNA is suspended in 25 μl of water.

### Phosphorylation of the vector

 $25\,\mu l$  of the vector prepared in the preceding step are diluted in a final volume of 500  $\mu l$  of the following reaction mixture:

After repair, the DNA is subjected to a phenol-chloroform extraction and a precipitation, the pellet is then taken up in 10 µl of water, the DNA is quantified by measuring the optical density at 260 nm. The quantified DNA is ligated into the vector PGEm-5Zf(+) prepared by the restriction

enzyme EcoRV and dephosphorylated (see preparation of the vector). The ligation is carried out under three conditions which vary in the ratio between the number of vector molecules and the number of insert molecules. Typically, an equimolar ratio, a ratio of 1:3 and a ratio of 3:1 are used for the ligations which are, moreover, carried out under the following conditions: vector PGEm-5Zf(+) 5 25 ng, cut DNA, ligation buffer in a final volume of 20 μl with T4 DNA ligase (Amersham E70042X); the whole is then placed in a refrigerator overnight and then a phenol-chloroform extraction and a precipitation are carried out in a conventional manner. The pellet is taken up in 5 µl of water.

### Transformation of the bacteria

### Plating of the bacteria

Petri dishes containing LB Agar medium containing ampicillin (50 µg/ml), Xgal [5-bromo-4-chloro-indolyl-beta-D-galactopyranoside (Sigma B-4252)], **IPTG**  $(280 \mu g/ml)$ (140 µg/ml) [isopropyl-beta-D-thiogalactoside (Sigma I-6758)] are used, 50 and 100 µl of bacteria are plated for each of the ligations. The Petri dishes are placed upside down at 37°C for 15 to 16 hours in an oven. The number of "recombinant" positive clones is evaluated by counting the white colonies and 15 the blue colonies which are thought to contain the vector alone.

### Evaluation of the "recombinant" positive clones

Ninety-four white colonies and two blue colonies are collected with the aid of sterile cones and are deposited at the bottom of the wells of plates designed for carrying out the amplification techniques. 30 µl of the following reaction mixture are added to each well: 1.7 mM MgCl<sub>2</sub>, 0.2 mM 20 each of dATP, dCTP, dGTP and dTTP, two synthetic oligonucleotides corresponding to sequences flanking the cloning site on either side and orienting the synthesis of the DNA in a convergent manner (0.5 µM RP and PU primers, 1 U TAQ polymerase (GibcoBRL 18038-026)).

The colonies thus prepared are subjected to a temperature of 94°C for 5 min and then to 30 thermal cycles composed of the following steps: 94°C for 40 s, 50°C for 30 s, 72°C for 180 s. The 25 reaction is then kept for 7 min at 72°C and then kept at 4°C.

The amplification products are deposited on an agarose gel (0.8%), stained with ethidium bromide, subjected to electrophoresis, and then analysed on an ultraviolet table. The presence of an amplification fragment having a size greater than 500 base pairs indicates the presence of an insert. The bacterial clones are then prepared so as to study the sequence of their insert.

#### 30 Sequencing

To sequence the inserts of the clones obtained as above, these were amplified by PCR on bacteria cultures carried out overnight using the primers for the vectors flanking the inserts. The sequence of the ends of these inserts (on average 500 bases on each side) was determined by automated fluorescent sequencing on an ABI 377 sequencer, equipped with the ABI Prism DNA 35 Sequencing Analysis software (version 2.1.2).

### Analysis of the sequences

by sequencing in a high-yield line (Figure 1) are obtained The sequences stored in a database; this part of the production is independent of any treatment of the sequences. The sequences are extracted from the database, avoiding all the regions of inadequate quality, that is to say the regions for which uncertainties are observed on the sequence at more than 95%. After extraction, 5 the sequences are introduced into a processing line, the diagram of which is described in Figure 2. In a first path of this processing line, the sequences are assembled by the Gap4 software from R. Staden (Bonfield et al., 1995) (OS UNIX/SUN Solaris); the results obtained by this software are kept in the form of two files which will be used for a subsequent processing. The first of these files provides information on the sequence of each of the contigs obtained. The second file represents all the clones 10 participating in the composition of all the contigs as well as their positions on the respective contigs.

The second processing path uses a sequence assembler (TIGR-Asmg assembler UNIX/SUN Solaris); the results of this second processing path are kept in the form of a file in the TIGR-Asmg format which provides information on the relationship existing between the sequences selected for the assembly. This assembler is sometimes incapable of linking contigs whose ends 15 overlap over several hundreds of base pairs.

The results obtained from these two assemblers are compared with the aid of the BLAST program, each of the contigs derived from one assembly path being compared with the contigs derived from the other path.

For the two processing paths, the strict assembly parameters are fixed (95% homology, 20 30 superposition nucleotides). These parameters avoid 3 to 5% of the clones derived from eukaryotic cells being confused with sequences obtained from the clones derived from Chlamydia pneumoniae. The eukaryotic sequences are however preserved during the course of this project; the strategy introduced, which is described below, will be designed, inter alia, not to be impeded by these sequences derived from contaminating clones.

The results of these two assemblers are processed in a software developed for this project. This software operates on a Windows NT platform and receives, as data, the results derived from the STADEN software and/or the results derived from the TIGR-Asmg assembler, the software, results, after processing of the data, in the determination of an assembly map which gives the proximity relationship and the orientation of the contigs in relation to one another (Figure 3a). Using 30 this assembly map, the software determines all the primers necessary for finishing the project. This treatment, which will be detailed below, has the advantage of distinguishing the isolated sequences derived from the contaminations, by the DNA eukaryotic cells, of the small-sized sequences clearly integrated into the project by the relationships which they establish with contigs. In order to allow, without any risk of error, the arrangement and the orientation of the contigs in relation to one another, 35 a statistical evaluation of the accuracy of the names (naming) "naming" of sequence is made from the results of "contigation". This evaluation makes it possible to give each of the clone plates, as well as each of the subsets of plates, a weight which is inversely proportional to probable error rate existing in the "naming" of the sequences obtained from this plate or from a subset of this plate. In spite of a low error rate, errors may occur throughout the steps of production of the clones and of the sequences. These steps are numerous, repetitive and although most of them are automated, others, like the deposition in the sequencers, are manual; it is then possible for the operator to make mistakes such as the inversion of two sequences. This type of error has a repercussion on the subsequent processing of the data, by resulting in relationships (between the contigs) which do not exist in reality, then in attempts at directed sequencing between the contigs which will end in failure. It is because of this that the evaluation of the naming errors is of particular importance since it allows the establishment of a probabilistic assembly map from which it becomes possible to determine all the clones which will serve as template to obtain sequences separating two adjacent contigs. Table 2 of parent U.S. application serial No. 60/107078 filed November 4, 1998 and French application 97-14673 filed November 21, 1997, each of which is incorporated by reference herein in its entirety, gives the clones and the sequences of the primers initially used during the initial operations.

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To avoid the step which consists in ordering and then preparing the clones by 15 conventional microbiological means, outer and inner primers oriented towards the regions not yet sequenced are defined by the software. The primers thus determined make it possible to prepare, by PCR, a template covering the nonsequenced region. It is the so-called outer primers (the ones most distant from the region to be sequenced) which are used to prepare this template. The template is then purified and a sequence is obtained on each of the two strands during 2 sequencing reactions which 20 each use one of the 2 inner primers. In order to facilitate the use of this approach, the two outer primers and the two inner primers are prepared and then stored on the same position of 4 different 96well plates. The two plates containing the outer primers are used to perform the PCRs which will serve to prepare the templates. These templates will be purified on purification columns preserving the topography of the plates. Each of the sequences will be obtained using primers situated on one and 25 then on the other of the plates containing the inner primers. This distribution allows a very extensive automation of the process and results in a method which is simple to use for finishing the regions not yet sequenced. Table 3 of parent U.S. application serial No. 60/107078 filed November 4, 1998 and French application 97-14673 filed November 21, 1997, each of which is incorporated by reference herein in its entirety, gives the names and the sequences of the primers used for finishing Chlamydia 30 pneumoniae.

Finally, a number of contigs exist in a configuration where one of their ends is not linked to any other contig end (Figure 3b) by a connecting clone relationship (a connecting clone is defined as a clone having one sequence end on a contig and the other end of its sequence on another contig; furthermore, this clone must be derived from a plate or a subset of plates with adequate naming quality). For the *Chlamydia pneumoniae* project, this particular case occurred 24 times. Two adjacent PCR primers orienting the synthesis of the DNA towards the end of the consensus sequence are defined for each of the orphan ends of the consensus sequence. The primer which is closest to the end

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of the sequence is called the inner primer whereas the primer which is more distant from the end of the sequence is called the outer primer. The outer primers are used to explore the mutual relationship between the orphan ends of the different contigs. The presence of a single PCR product and the possibility of amplifying this product unambiguously using the inner primers evokes the probable 5 relationship between the contigs on which the primers which allowed the amplification are situated. This relationship will be confirmed by sequencing and will allow the connection between the orphan ends of the consensus sequences. This strategy has made it possible to obtain a complete map of the Chlamydia pneumoniae chromosome and then to finish the project.

### **Quality control**

All the bases not determined with certainty in the chromosomal sequence were noted and the density of uncertainties was measured on the entire chromosome. The regions with a high density of uncertainties were noted and the PCR primers spanning these regions were drawn and are represented in Table 4 of parent U.S. application serial No. 60/107078 filed November 4, 1998 and French application 97-14673 filed November 21, 1997 each of which is incorporated by reference 15 herein in its entirety.

The sequence of each of the PCR products was obtained with two operational primers different from the amplification primers. The sequences were obtained in both directions for all the PCRs (100% success).

### Data banks

Local reorganizations of major public banks were used. The protein bank used consists of the nonredundant fusion of the Genpept bank (automated translation of GenBank, NCBI; Benson et al., 1996).

The entire BLAST software (public domain, Altschul et al., 1990) for searching for homologies between a sequence and protein or nucleic data banks was used. The significance levels 25 used depend on the length and the complexity of the region tested as well as the size of the reference bank. They were adjusted and adapted to each analysis.

The results of the search for homologies between a sequence according to the invention and protein or nucleic data banks are presented and summarized in Table 1 below.

# 30 Table 1: List of coding chromosome regions and homologies between these regions and the sequence banks.

Legend to Table 1: Open reading frames are identified with the GenMark software version 2.3A (GenePro), the template used is Chlamydia pneumoniae of order 4 on a length of 196 nucleotides with a window of 12 nucleotides and a minimum signal of 0.5. The reading frames 35 ORF2 to ORF 1137 are numbered in order of appearance on the chromosome, starting with ORF2 (ORF column). The positions of the beginning and of the end are then given in column 2 (position). When the position of the beginning is greater than the position of the end, this means that the region is

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encoded by the strand complementary to the sequence which was given in the sequence SEQ ID No. 1.

All the putative products were subjected to a search for homology on GENPEPT (release 102 for SEQ ID No. 2 to SEQ ID No. 1137, and release 108 for SEQ ID No. 1138 to SEQ ID No. 1291 and SEQ ID No. 6844 to SEQ ID No. 6849) with the BLASTP software (Altschul et al. 1990). With, as parameters, the default parameters with the exception of the expected value E set at 10<sup>-5</sup> (for SEQ ID No. 2 to SEQ ID No. 1137) and P value set at e<sup>-10</sup> (for SEQ ID No. 1138 to SEQ ID No. 1291 and SEQ ID No. 6844 to SEQ ID No. 6849). Subsequently, only the identities greater than 30% (I% column) were taken into account. The description of the most homologous sequence is given in the Homology column; the identifier for the latter sequence is given in the ID column and the animal species to which this sequence belongs is given in the Species column. The Homology score is evaluated by the sum of the blast scores for each region of homology and reported in the Score column.

### Materials and Methods for transmembrane domains:

The DAS software was used as recommended by the authors (Cserzo et al., 1997).

This method uses, to predict the transmembrane domains, templates derived from a sampling of selected proteins. All the regions for which a "Cutoff" greater than 1.5 was found by the program were taken into account.

### Additional ORF Finder Programs

For this analysis, two additional ORF finder programs were used to predict potential open reading frames of a minimum length of 74 amino acids; Glimmer (Salzberg, S.L., Delcher, A., Kasif, S., and W. White. 1998. Microbial gene identification using interpolated Markov models. Nucleic Acids Res. 26:544-548.), and an in-house written program. The in-house program used a very simple search algorithm. The analysis required the that the genomic DNA sequence text be in the 5' to 3' direction, the genome is circular, and that TAA, TAG, and TGA are stop codons. The search parameters were as follows:

- (1) A search for an ORF that started with a GTG codon was performed. If no GTG codons were found, then a search for an ATG codon was performed. However, if a GTG codon was found, then a search downstream for a ATG codon was performed. All start and stop nucleotide positions were recorded.
- (2) A search for an ORF that started with a TTG codon was performed. If no TTG codons were found, then a search for a ATG codon was performed. However, if a TTG codon was found, then a search downstream for a ATG codon was performed. All start and stop nucleotide positions were recorded.
- (3) The analysis described in steps 1 and 2 were repeated for the opposite strand of DNA sequence.

- (4) A search for ORFs that determined all ORF lengths using start and stop positions in the same reading frames was performed.
- (5) All ORFs whose DNA length was less than 225 nucleotides were eliminated from the search.

### 5 Surface Exposed Protein Search Criteria

Potential cell surface vaccine targets are outer membrane proteins such as porins, lipoproteins, adhesions and other non-integral proteins. In *Chlamydia psittaci*, the major immunogens is a group of putative outer membrane proteins (POMPs) and no homologs have been found in *Chlamydia pneumoniae* and *Chlamydia trachomatis* by traditional analysis (Longbottom, D., Russell, M., Dunbar, S.M., Jones, G.E., and A.J. Herring. 1998. Molecular Cloning and Characterization of the Genes Coding for the Highly Immunogenic Cluster of 90-Kilodalton Envelope Proteins from *Chlamydia psittaci* Subtype That Causes Abortion in Sheep. Infect Immun 66:1317-1324.) Several putative outer membrane proteins have been identified in *Chlamydia pneumoniae*, all of which may represent vaccine candidates. The major outer membrane protein (MOMP) gene (omp1) has been found in various isolates of *Chlamydia pneumoniae* (Jantos, CA., Heck, S., Roggendorf, R., Sen-Gupta, M., and Hegemann, JH. 1997. Antigenic and molecular analyses of different chlamydia pneumoniae strains. J. Clin Microbiology 35(3):620-623.) Various criteria, as listed below, were used to identify putative surface exposed ORFs from the genomic DNA sequence of *Chlamydia pneumoniae* (French application 97-14673 filed 21 November 1997). Any ORF which met any one or more of the individual criteria were listed in this category.

Protein homology searches were done using the Blastp 2.0 tool (Altschul, S.F., Madden, T.L., Schaffer, A.A., Zhang, J., Zhang, Z., Miller, W., and D.J. Lipman. 1997. Gapped BLAST and PSI-BLAST: a new generation of protein database search programs. Nucleic Acids Res. 25:3389-3402.) An ORF product was labeled surface exposed if there was homology to a known, or hypothetical, or putative surface exposed protein with a P score better than e<sup>-10</sup>.

Most, if not all, proteins that are localized to the membrane of bacteria, via a secretory pathway, contain a signal peptide. A software program, SignalP, analyzes the amino acid sequence of an ORF for such a signal peptide (Nielsen, H., Engelbrecht. J., Brunak, S., and G. von Heijne. 1997. Identification of prokaryotic and eukaryotic signal peptides and prediction of their cleavage sites.

Protein Engineering 10:1-6.) The first 60 N-terminal amino acids of each ORF were analyzed by SignalP using the Gram-Negative software database. The output generates four separate values, maximum C, maximum Y, maximum S, and mean S. The S-score, or signal region, is the probability of the position belonging to the signal peptide. The C-score, or cleavage site, is the probability of the position being the first in the mature protein. The Y-score is the geometric average of the C-score and a smoothed derivative of the S-score. A conclusion of either a Yes or No is given next to each score. If all four conclusions are Yes and the C-terminal amino acid is either a phenylalanine (F) or a tyrosine (Y), the ORF product was labelled outer membrane (Struyve, M., Moons, M., and J. Tommassen.

1991. Carboxy-terminal Phenylalanine is Essential for the Correct Assembly of a Bacterial Outer Membrane Protein. J. Mol. Biol. 218:141-148.)

The program called Psort, determines the localization of a protein based on its signal sequence, recognition of transmembrane segments, and analysis of its amino acid composition (Nakai, 5 K., and M. Kanehisa. 1991. Expert system for predicting protein localization sites in gram-negative bacteria. Proteins 11:95-110.) An ORF product is considered to be an outer membrane protein if the output data predicts the protein as outer membrane with a certainty value of 0.5 or better and whose value is at least twice as large as the next predicted localized certainty value.

Finally, ORF products that were not predicted to be outer membrane or surface exposed,

10 based on the above criteria, were further analyzed. The blastp output data for these ORFs were
searched using various general and specific keywords, suggestive of known cell surface exposed
proteins. An ORF was labeled surface exposed if the keywords matched had a Blastp hit, had a P
score better than e<sup>-10</sup>, and that there was no better data indicating otherwise. The following is a list of
the searched keywords:

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	Adhesion	Adhesin	Invasin	Invasion	Extensin	
	Omp	Outer Surface	Porin	Outer Membra	ne	
	Cell Surface	Cell Wall	Pilus	Pilin	Flagellar sheath	BtuB
	Cir	ChuA	CopB	ExeD	FadL	FecA
20	FepA	FhuA	FmdC	FomA	FrpB	GspD
	HemR	HgbA	Hgp	HmbR	HmuR	HMW
	HrcC	Hrp	InvG	LamB	LbpA	LcrQ
	Lmp1	MxiD	MOMP	PilE	HpaA	NolW
	NspA	OpcP	OpnP	Opr	OspA	PhoE
25	PldA	Por	PscC	PulD	PupA	QuiX
	RafY	ScrY	SepC	ShuA	SomA	SpiA
	Tbp1	Yop	YscC	mip	Tol	

Those ORFs that did not meet the minimum requirement for being an outer membrane protein based on the above search criteria but which were homologous to identified outer membrane ORFs in Chlamydia trachomatis were included. The Chlamydia trachomatis genome (French patent applications FR97-15041, filed 28 November 1997 and 97-16034 filed 17 December 1997) was analyzed using the above search criteria and a number of outer membrane ORFs were identified. These Chlamydia trachomatis ORFs were then tested against the Chlamydia pneumoniae genome using Blastp. Any Chlamydia pneumoniae ORF with a Blastp P value better than e-10 against a Chlamydia trachomatis outer membrane was included in this section, if there was no better data

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A list of ORFs in the Chlamydia pneumoniae genome encoding indicating otherwise. putative surface exposed proteins is set forth above in the specification.

# Identification of Putative Lipoproteins in the Genome of Chlamydia pneumoniae

Lipoproteins are the most abundant post-translationally modified bacterial secretory proteins (Pugsley, A. P., 1993. The complete general secretory pathway in Gramnegative bacteria. Microbiol. Rev. 57:50-108). The characteristic features of lipoproteins are a thiol-linked diacylglyceride and an amine-linked monoacyl group on the cysteine that becomes the amino-terminal residue after signal peptide cleavage by Signal Peptidase II. 10 (Pugsley, A. P.. 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108). The identification of putative lipoproteins from the genomic sequencing of Chlamydia pneumoniae was done by examining the deduced amino acid sequence of identified ORFs for the presence of a signal peptide with a Signal Peptidase II cleavage site analogous to the consensus sequence for prolipoprotein modification and 15 processing reactions (Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York; Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria. J. Bacteriol. 177:1123-1128.).

Chlamydia pneumoniae ORFs were initially screened for the most basic of lipoprotein characteristics, a cysteine in the first 30 amino acids of the deduced protein. ORFs with a standard start codon (ATG, GTG, or TTG) and having one or more of the following characteristics were selected for direct analysis of their first 30 amino acids:

- (a) Significant Signal P value (at least two out of the four values are Yes)
- (b) PSORT value indicating membrane passage (IM-inner membrane, Peri-periplasm, or OM-outer membrane)
- (c) Identification of the word lipoprotein among the ORF blastp data set.
- (d) A Blastp value of <e<sup>-10</sup> with a putative lipoprotein from Chlamydia trachomatis 30 (French applications 97-15041 filed 28 November 1997 and 97-16034 filed 17 December 1997).

The first 30 amino acids of each ORF in this set were analyzed for the characteristics commonly found in lipoprotein signal peptides (Pugsley, A. P., 1993. The complete general secretory 35 pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108; Hayashi, S., and H. C. Wu. 1992.

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Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York; Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria, J. Bacteriol. 177:1123-1128.) Putative lipoprotein signal peptides were required to have a 5 cysteine between amino acid 10 and 30 and reach a minimum score of three based on the following criteria for lipoprotein signal peptides:

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- (a) Identification of specific amino acids in specific positions around the cysteine which are part of the consensus Signal Peptidase II cleavage site (Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York); Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria. J. Bacteriol. 177:1123-1128). Since the identification of the cleavage site is the most important factor in identifying putative lipoproteins, each correctly positioned amino acid contributed toward reaching the minimum score of three. (b) A hydrophobic region rich in alanine and leucine prior to the cleavage site (Pugsley, A. P.. 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108) contributed toward reaching the minimum score of three.
  - (c) A short stretch of hydrophilic amino acids greater than or equal to 1 usually lysine or arginine following the N-terminal methionine (Pugsley, A. P.. 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108) contributed toward reaching the minimum score of three.

A list of ORFs in the Chlamydia pneumoniae genome encoding putative lipoproteins is set forth above in the specification.

#### LPS-Related ORFs of Chlamydia pneumoniae 25

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Lipopolysaccharide (LPS) is an important major surface antigen of Chlamydia cells. Monoclonal antibodies (Mab) directed against LPS of Chlamydia pneumoniae have been identified that can neutralize the infectivity of Chlamydia pneumoniae both in vitro and in vivo (Peterson, E.M., de la Maza, L.M., Brade, L., Brade, H. 1998. Characterization of a Neutralizing Monoclonal 30 Antibody Directed at the Lipopolysaccharide of Chlamydia pneumonia. Infect. Immun. Aug. 66(8):3848-3855.) Chlamydial LPS is composed of lipid A and a core oligosaccharide portion and is phenotypically of the rough type (R-LPS) (Lukacova, M., Baumann, M., Brade, L., Mamat, U., Brade, H. 1994. Lipopolysaccharide Smooth-Rough Phase Variation in Bacteria of the Genus Chlamydia. Infect. Immun. June 62(6):2270-2276.) The lipid A component is composed of fatty acids 35 which serve to anchor LPS in the outer membrane. The core component contains sugars and sugar derivatives such as a trisaccharide of 3-deoxy-D-manno-octulosonic acid (KDO) (Reeves, P.R., Hobbs, M., Valvano, M.A., Skurnik, M., Whitfield, C., Coplin, D., Kido, N., Klena, J., Maskell, D.,

Raetz, C.R.H., Rick, P.D. 1996. Bacterial Polysaccharide and Gene Synthesis Nomenclature pp. 10071-10078, Elsevier Science Ltd.). The KDO gene product is a multifunctional glycosyltransferase and represents a shared epitope among the Chlamydia. For a review of LPS biosynthesis see, e.g., Schnaitman, C.A., Klena, J.D. 1993. Genetics of Lipopolysaccharide 5 Biosynthesis in Enteric Bacteria. Microbiol. Rev. 57:655-682.

A text search of the ORF blastp results identified several genes that are involved in Chlamydial LPS production with a P score better than e<sup>-10</sup>. The following key-terms were used in the text search: KDO, CPS (Capsular Polysaccharide Biosynthesis), capsule, LPS, rfa, rfb, rfc, rfe, rha, rhl, core, epimerase, isomerase, transferase, pyrophosphorylase, phosphatase, aldolase, heptose, 10 manno, glucose, lpxB, fibronectin, fibrinogen, fucosyltransferase, lic, lgt, pgm, tolC, rol, ChoP, phosphorylcholine, waaF, PGL-Tb1. A list of ORFs in the Chlamydia pneumoniae genome encoding putative polypeptides involved in LPS biosynthesis is set forth above in the specification.

### Type III And Other Secreted Products

Type III secretion enables gram-negative bacteria to secrete and inject pathogenicity proteins into the cytosol of eukaryotic host cells (Hueck, C. J., 1998. Type III Protein Secretion Systems in Bacterial Pathogens of Animals and Plants. In Microbiology and Molecular Biology Reviews. 62:379-433.) These secreted factors often resemble eukaryotic signal transduction factors, thus enabling the bacterium to redirect host cell functions (Lee, C.A., 1997. Type III secretion 20 systems: machines to deliver bacterial proteins into eukaryotic cells? Trends Microbiol. 5:148-156.) In an attempt to corrupt normal cellular functions, Chlamydial pathogenicity factors injected into the host cytosol will nonetheless, as cytoplasmic constituents be processed and presented in the context of the Major Histocompatibility Complex (MHC class I). As such, these pathogenicity proteins represent MHC class I antigens and will play an important role in cellular immunity. Also included in this set are secreted non-type III products that may play a role as vaccine components.

A text search of the ORF blastp results identified genes that are involved in Chlamydia pneumoniae protein secretion with a P score better than e<sup>-10</sup>. The following key-terms were used in the text search in an effort to identify surface localized or secreted products: Yop, Lcr, Ypk, Exo, Pcr, Pop, Ipa, Vir, Ssp, Spt, Esp, Tir, Hrp, Mxi, hemolysin, toxin, IgA protease, cytolysin, tox, hap, 30 secreted and Mip.

Chlamydia pneumoniae ORFs that did not meet the above keyword search criteria, but have homologs in Chlamydia trachomatis that do meet the search criteria are included herein. The Chlamydia trachomatis genome (French patent applications FR97-15041, filed 28 November 1997 and 97-16034 filed 17 December 1997) was analyzed using the above search criteria and a number of 35 ORFs were identified. These Chlamydia trachomatis ORFs were tested against the Chlamydia pneumoniae genome using Blastp. Any Chlamydia pneumoniae ORF with a Blastp P value < e-10 against a Chlamydia trachomatis homolog, identified using the above search criteria, was included. A

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list of ORFs in the Chlamydia pneumoniae genome encoding putative secreted proteins is in the specification.

# Chlamydia pneumoniae: RGD Recognition Sequence

Proteins that contain Arg-Gly-Asp (RGD) attachment site, together with integrins that serve as their receptor constitute a major recognition system for cell adhesion. The RGD sequence is the cell attachment site of a large number of adhesive extracellular matrix, blood, and cell surface proteins and nearly half of the known integrins recognize this sequence in their adhesion protein ligands. There are many RGD containing microbial proteins such as the penton protein of adenovirus, 10 the coxsackie virus, the foot and mouth virus and pertactin, a 69 kDa (kilodalton) surface protein of Bordetella pertussis, that serve as ligands through which these microbes bind to integrins on the cell surfaces and gain entry into the cell. The following provides evidence supporting the importance of RGD in microbial adhesion:

- a) The adenovirus penton base protein has a cell rounding activity and when penton base was expressed in E. coli, it caused cell rounding and cells adhered to polystyrene wells coated with the protein. Mutant analysis showed that both these properties required an RGD sequence. Virus mutants with amino acid substitutions in the RGD sequence, showed much less adherence to HeLa S3 cells, and also were delayed in virus reproduction (Bai, M., Harfe, B., and Freimuth, P. 1993. Mutations That Alter an RGD Sequence in the Adenovirus Type 2 Penton Base Protein Abolish Its Cell-Rounding Activity and Delay Virus Reproduction in Flat Cells. J. Virol. 67:5198-5205).
- b) It has been shown that attachment and entry of coxsackie virus A9 to GMK cells were dependent on an RGD motif in the capsid protein VP1. VP1 has also been shown to bind  $\alpha_v \beta_3$ integrin, which is a vitronectin receptor (Roivainen, M., Piirainen, L., Hovi, T., Virtanen, I., Rijkonen, T., Heino, J., and Hyypia, T. 1994. Entry of Coxsackievirus A9 into Host Cells: Specific Interactions with a,b, Integrin, the Vitronectin Receptor Virology, 203:357-65).
- c) During the course of whooping cough, Bordetella pertussis interacts with alveolar macrophages and other leukocytes on the respiratory epithelium. Whole bacteria adheres by means of two proteins, filamentous hemagglutinin (FHA) and pertussis toxin. FHA interacts with two classes of molecules on macrophages, galactose containing glycoconjugates and the integrin CR3. The interaction between CR3 and FHA involves recognition of RGD sequence at the positions 1097-1099 in FHA (Relman, D., Tuomanen, E., Falkow, S., Golenbock, D. T., Saukkonen, K., and Wright, S. D. "Recognitition of a Bacterial Adhesin by an Integrin: Macrophage CR3 Binds Filamentous Hemagglutinin of Bordetella Pertussis." Cell, 61:1375-1382 (1990)).

- d) Pertactin, a 69 kDa outer membrane protein of *Bordetella pertussis*, has been shown to promote attachment of Chinese hamster ovary cells (CHO). This attachment is mediated by recognition of RGD sequence in pertactin by integrins on CHO cells and can be inhibited by synthetic RGD containing peptide homologous to the one present in pertactin (Leininger, E., Roberts, M., Kenimer, J. G., Charles, I. G., Fairweather, N., Novotny, P., and Brennan, M. J. 1991. Pertactin, an Arg-Gly-Asp containing *Bordetella pertussis* surface protein that promotes adherence of mammalian cells Proc. Natl. Acad. Sci. USA, 88:345-349).
- e) The RGD sequence is highly conserved in the VP1 protein of foot and mouth disease virus (FMDV). Attachment of FMDV to baby hamster kidney cells (BHK) has been shown to be mediated by VP1 protein via the RGD sequence. Antibodies against the RGD sequence of VP1 blocked attachment of virus to BHK cells (Fox, G., Parry, N. R., Barnett, P. V., McGinn, B., Rowland, D. J., and Brown, F. 1989. The Cell Attachment Site on Foot-and-Mouth Disease Virus Includes the Amino Acid Sequence RGD (Arginine-Glycine-Aspartic Acid) J. Gen. Virol., 70:625-637).

It has been demonstrated that bacterial adherence can be based on interaction of a bacterial adhesin RGD sequence with an integrin and that bacterial adhesins can have multiple binding site characteristic of eukaryotic extracellular matrix proteins. RGD recognition is one of the important mechanisms used by microbes to gain entry into eukaryotic cells.

The complete deduced protein sequence of the Chlamydia pneumoniae genome was searched for the presence of RGD sequence. There were a total of 54 ORFs that had one or more RGD sequences. Not all RGD containing proteins mediate cell attachment. It has been shown that RGD containing peptides that have proline immediately following the RGD sequence are inactive in cell attachment assays (Pierschbacher & Ruoslahti. 1987. Influence of stereochemistry of the sequence Arg-Gly-Asp-Xaa on binding specificity in cell adhesion. J. Biol. Chem. 262:17294-98). ORFs that had RGD, with proline as the amino acid following the RGD sequence were excluded from the list. Also, RGD sequence may not be available at the surface of the protein or may be present in a context that is not compatible with integrin binding. Since not all RGD- containing proteins are involved in cell attachment, several other criteria were used to refine the list of RGD- containing proteins. A list of ORFs in the Chlamydia pneumoniae genome encoding polypeptides with RGD recognition sequence(s) is in the specification.

### Non-Chlamydia trachomatis ORFs

35 Chlamydia pneumoniae ORFs were compared to the ORFs in the Chlamydia trachomatis genome (French patent applications FR97-15041, filed 28 November 1997 and 97-16034 filed 17 December 1997) using Blastp. Any Chlamydia pneumoniae ORF with a Blastp P value worse than e

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<sup>10</sup> (i.e. >e<sup>-10</sup>) against *Chlamydia trachomatis* ORFs are included in this section. A list of ORFs in the *Chlamydia pneumoniae* genome which are not found in *Chlamydia trachomatis* is set forth above in the specification.

### Cell Wall Anchor Surface ORFs

Many surface proteins are anchored to the cell wall of Gram-positive bacteria via the conserved LPXTG motif (Schneewind, O., Fowler, A., and Faull, K.F. 1995. Structure of the Cell Wall Anchor of Surface Proteins in Staphylococcus aureus. Science 268:103-106). A search of the Chlamydia pneumoniae ORFs was done using the motif LPXTG. A list of ORFs in the Chlamydia pneumoniae genome encoding polypeptides anchored to the cell wall is in the specification.

### **ATCC Deposits**

Samples of Chlamydia pneumoniae were deposited with the American Type Culture Collection (ATCC), Rockville, Maryland, on November 19, 1998 and assigned the accession number ---. Cells can be grown, harvested and purified, and DNA can be prepared as discussed above. In order to enable recovery of specific fragments of the chromosome, one can run targeted PCR reactions, whose amplification products can then be sequenced and/or cloned into any suitable vector, according to standard procedures known to those skilled in the art.

In addition, a sample of three pools of clones covering chromosomal regions of interest were deposited with the American Type Culture Collection (ATCC), Rockville, Maryland, on November 19, 1998 and assigned the indicated accession number: — . Each pool of clones contains a series of clones. When taken together, the three pools in the sample cover a portion of the chromosome, with a redundancy of slightly more than two. The total number of clones in the sample is 196.

The clones cover the following three regions of interest:

- (i) position 30,000 to 40,000 of SEQ ID No. 1, referred to as region A;
- (ii) position 501,500 to 557,000 of SEQ ID No. 1, referred to as region B; and
- (iii) position 815,000 to 830,000 of SEQ ID No. 1, referred to as region C.

Table 4 lists groups of oligonucleotides to be used to amplify each of ORFs 2-1291 according to standard procedures known to those skilled in the art. Such oligonucleotides are listed as SEQ ID Nos. 1292 to 6451. For each ORF, the following is listed: one forward primer positioned 2,000 bp upstream of the beginning of the ORF; one forward primer positioned 200 bp upstream of the beginning of the ORF; one reverse primer positioned 2,000 bp downstream at the end of ORF, which is 2,000 bp upstream of the end site of the ORF on the complementary strand; and one reverse primer 200 bp downstream at the end of ORF, which is 200 bp upstream of the end site of the ORF on the complementary strand. The corresponding SEQ ID Nos. for the primers are listed in Table 4, where Fp is the proximal forward primer; Fd is the distal forward

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primer; Bp is the proximal reverse primer; and Bd is the distal reverse primer. The positions of the 5' ends of each of these primers on the nucleotide sequence of SEQ ID No. 1 are shown in Table 5.

Table 6 lists oligonucleotides (SEQ ID Nos. 6452-6843) to be used to amplify the inserts of each of the 196 clones present in the pooled sample according to standard procedures well known to those of skill in the art. These primers can also be utilized to amplify the chromosomal region corresponding to the region A, B or C within which the particular insert lies. Their positions are indicated in Table 7.

The present invention is not to be limited in scope by the specific embodiments described herein, which are intended as single illustrations of individual aspects of the invention, and functionally equivalent methods and components are within the scope of the invention. Indeed, various modifications of the invention, in addition to those shown and described herein will become apparent to those skilled in the art from the foregoing description and accompanying drawings. Such modifications are intended to fall within the scope of the appended claims.

All publications and patent applications mentioned in this specification are herein incorporated by reference to the same extent as if each individual publication or patent application was specifically and individually indicated to be incorporated by reference.

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			TARIE 1				
ORF	Begin	End	Homology	QI	Species	Score	%1
ORF2	42	794	triosephosphate isomerase	L27492	Thermotoga maritima	292	54
ORF3	1258	1614	putative			;	,
ORF4	1807	2418	polypeptide deformylase	D90906	Synechocystis sp.	310	9 5
ORFS	3393	2491	hypothetical protein	Z75208	Bacillus subtilis	338	75
ORF6	3639	4067	unknown	U87792	Bacillus subtilis		28
ORF7	5649	4270	putative				
ORF8	7463	6012	putative				T
ORF9	8051	8962	putative				$\top$
ORF10	9129	6566	putative				T
ORF11	10687	10361	putative				
ORF12	10927	11232	putative				Ç
ORF13	11246	12727	amidase	U49269	Moraxella catarrhalis	2011	74
ORF14	12691	14190	PET112	D90913	Synechocystis sp.	1044	9
ORF15	14484	17249	POMP91A	U65942	Chlamydia psittaci	10/4	543
ORF16	16039	15770	putative				
ORF17	17845	20853	putative				
ORF18	21137	22042	putative				
ORF19	22046	23476	putative				T
ORF20	23681	26110	putative				T
ORF21	26109	25861	putative				
ORF22	26241	26978	putative				
ORF23	26960	27754	putative				
ORF24	27747	28577	putative			60,	6
ORF25	28887	29492	POMP91A	U65942	Chlamydia psittaci	180	3
ORF26	29432	30028	POMP91A	U65942	Chlamydia psittaci	301	7
ORF27	30024	31472	POMP91A	U65942	Chlamydia psittaci	8/9	4
ORF28	31758	32288	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	4	43
ORF29	32201	33991	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	1126	84 (
ORF30	33852	34541	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	589	3
ORF31	34783	36063	POMP91B precursor	U65943	Chlamydia psittaci	469	<u>و</u> :
ORF32	36009	37529	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	1338	7
ORF33	37881	39362	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	1/9	94

ORF	Begin	End	Homology	Œ	Species	Score	<u>~</u>
ORF34	39418	39161	putative			H	
ORF35	39366	40715	POMP90A precursor	U65942	Chlamydia psittaci	904	42
ORF36	43076	41094	putative				T
ORF37	43800	43066	putative				
ORF38	44828	43785	putative			$\dashv$	
ORF39	45340	44753	homologous to unidentified E. coli protein	M96343	Bacillus subtilis	-	44
ORF40	45752	45372	o530; This 530 aa orf is 33 pct identical (14	AE000184	Escherichia coli	769	43
			gaps) to 525 residues of an approx. 640 aa nrotein YHES HAEIN SW: P44808				
ORF41 .	46996	45701	ABC transporter, ATP-binding protein	AE000596	Helicobacter pylori	878	39.
			(yheS)				1
ORF42	47961	47569	putative				];
ORF43	48960	48040	hypothetical protein	D64001	Synechocystis sp.	404	37
ORF44	51452	50133	Lon protease-like protein	X74215	Homo sapiens	1232	54
ORF45	52606	51335	unknown	Z54285	Schizosaccharomyces pombe	781	44
ORF46	53684	53319	putative				$ \top $
ORF47	54195	53746	putative				
ORF48	55278	56453	heat-shock protein	U15010	Legionella pneumophila	975	45
ORF49	56493	57266	branched chain alpha-keto acid	M97391	Bacillus subtilis	329	36
			dehydrogenase E1-alpha				Ţ
ORF50	57297	58526	branched chain alpha-keto acid dehydrogenase E1-beta	M97391	Bacillus subtilis	707	20
ORFSI	59851	58565	putative		- 2		
ORF52	61495	59924	ComE	D90903	Synechocystis sp.	134	55
ORF53	61324	62151	putative		_		
54	62132	62470	Hpr protein	X12832	Bacillus subtilis	136	36
.55	62474	63733	enzyme I (ptsI)	U32844	Haemophilus influenzae	381	35
ORF56	63881	64186	f831; This 831 aa orf is 46 pct identical (11	AE000326	Escherichia coli	123	34
			gaps) to 709 residues of an approx. 712 aa				
_			protein PT1A ECOLI SW: P32670				
ORF57	64611	64318	ORF107	X17014	Bacillus subtilis	128	33
ORF58	65485	64673	putative				1
ORF59	62669	65301	dnaZX-like ORF put. DNA polymerase III	X06803	Bacillus subtilis	596	52

ORF	Begin	End	Homology	<b>a</b>	Species	Score	<u>%</u>
ORF60	66244	67281	putative				
ORF61	67265	66929	putative				T
ORF62	67703	68239	putative				
ORF63	68805	70736	putative				
ORF64	69172	68831	putative				
ORF65	70642	71142	putative				
ORF66	71325	72029	putative				
ORF67	72060	73637	putative			3	7
ORF68	74061	76175	YqfF	D84432	Bacillus subtilis	342	4 3
ORF69	78351	77680	porphobilinogen deaminase	D28503	Clostridium josui	262	42
ORF70	79356	78355	sms protein	D90914	Synechocystis sp.	736	22
ORF71	79983	79693	ribonuclease III (mc)	AE000579	Helicobacter pylori	88	33
ORF72	8041	79938	ORF3	D64116	Bacillus subtilis	268	4
ORF73	80475	69608	putative			- 1	[
OB F74	81296	83080	hypothetical protein	Y14079	Bacillus subtilis	893	38
OP E75	83201	83932	manganese superoxide dismutase	X77021	Caenorhabditis elegans	622	28
ORF76	84005	84769	acetyl-CoA carboxylase beta subunit (accD)	AE000604	Helicobacter pylori	602	20
00000	32000	7770	deoxymridinetrinhoenhatase (dut)	U32776	Haemophilus influenzae	110	41
ORF //	84973	95475	deoxonridine 5'-trinbosnhate	AE000596	Helicobacter pylori	265	89
\$ <del>5</del> 50	62169	02450	nucleotidohydrolase (dut)		- 22		
ORF79	85397	85903	ORF2	L26916	Pseudomonas aeruginosa	173	<u>%</u>
ORF80	85909	86583	enzyme IIANtr	U18997	Escherichia coli	2	47
ORF81	86626	88065	putative				
ORF82	89257	91026	putative		1		
ORF83	91291	93030	putative				
ORF84	93295	94086	putative		-		T
ORF85	95285	94707	putative				
ORF86	95667	96557	putative				T
ORF87	96317	97456	putative				
ORF88	98435	89626	putative				
ORF89	99460	98426	putative			1	2
ORF90	100144	101325	elongation factor Tu	L22216	Chlamydia trachomatis	191/	S

				_,	<del> ,</del>		- 1		_ ·	_	_				95		_			_					т		_		_	_		<del></del>	_
%1	9,	49	63	51	31	47	22	24	62	49					8	32	48		33	3	49	37	4	7	38			_	\$	8	<del>8</del>	_	
Score		376	458	642	82	325	2740	2947	514	755					934	147	751		264	184	1679	427	3107	1748	147				231	538	310		
Species		Thermus aquaticus thermophilus	Bacillus subtilis	Thermotoga maritima	Streptomyces antibioticus	Synechocystis PCC6803	Staphylococcus aureus	Escherichia coli	Leptospira biflexa	Homo sapiens					Thermus aquaticus thermophilus	Acetabularia acetabulum	Desulfurococcus sp. SY		Enterococcus hirae	Methanococcus jannaschii	Escherichia coli	Streptomyces coelicolor	Thermus thermophilus	Chlamydia trachomatis	Escherichia coli				Bacillus subtilis	Pseudomonas aeruginosa	Pseudomonas aeruginosa		11
<b>a</b>		L10348	D13303	Z11839	M89911	X53178	X64172	V00339	M22622	L19437					X63855	D50528	U96487		X76913	U67478	X05891	U19250	D49911	U83196	X61000				AB002150	Y10528	Y10528		
Homology	putative	transcription factor	ribosomal protein L11	ribosomal protein L1	ribosomal protein L10	rpl12 (AA 1-128)	DNA-directed RNA polymerase beta chain	rpoC	acetylomithine deacetylase (EC 5.1.1.16)	transaldolase	putative	putative	putative	putative	ATPase alpha-subunit	adenosine triphosphatase A subunit	V-ATPase B subunit	putative	v-type Na-ATPase	ATP synthase, subunit K	valyl-tRNA synthetase	protein kinase-like protein	UvrA	pyruvate kinase	HtrB protein	putative	putative	putative	YbbP	cyanide insensitive terminal oxidase	cyanide insensitive terminal oxidase	putative	putative
End	101720	102273	102805	103530	104058	104491	108386	112054	112590	113682	114121	114136	115755	116543	118055	118522	119843	120457	122430	122950	126347	129166	129213	136382	136482	138240	137928	138257	139516	141841	142658	143050	144494
Begin	101457	101704	102356	102835	103549	104096	104601	108401	112033	112672	113726	114711	115267	115911	116736	117968	118530	119816	120451	122504	123528	126332	134690	134925	137870	137899	138239	139558	140352	140498	141855	144258	145258
ORF	ORF91	ORF92	ORF93	ORF94	ORF95	ORF96	ORF97	ORF98	ORF99	ORF100	ORF101	ORF102	ORF103	ORF104	ORF105	ORF106	ORF107	ORF108	ORF109	ORF110	ORF111	ORF112	ORF113	ORF114	ORF115	ORF116	ORF117	ORF118	ORF119	ORF120	ORF121	ORF122	ORF123

ORF	Begin	End	Homology	αı	Species	Score	<u>~~~</u>
ORF124	145454	146749	product similar to E. coli PhoH protein	Z97025	Bacillus subtilis	836	47
ORF125	147318	146767	putative				
ORF126	148261	147677	putative				1
ORF127	149029	152157	isoleucyl-tRNA synthetase	U04953	Homo sapiens	2361	25
ORF128	154108	152201	leader peptidase I	D90904	Synechocystis sp.	225	47
ORF129	155135	154308	putative				!
ORF130	155141	155467	YtiA	AF008220	Bacillus subtilis	701	43
ORF131	155703	156779	orf 361; ranslated orf similarity to SW:	6968LX	Coxiella burnetii	863	- 65
			RF1_SALTY peptide chain release factor 1				
			of Salmonella typhimurium			-	
ORF132	156748	157635	product similar to E.coli PRFA2 protein	Z49782	Bacillus subtilis	144	37
ORF133	157653	158996	Ffh	U82109	Thermus aquaticus	797	45
ORF134	159363	159986	tRNA (guanine-N1)-methyltransferase	U32705	Haemophilus influenzae	545	49
			(trmD)				
ORF135	159880	160446	putative		1		
ORF136	160477	160839	ribosomal protein L19	X72627	Synechocystis sp.	319	တ္က
ORF137	160898	161539	putative protein highly homologous to E.	D32253	Magnetospirillum sp.	427	49
			coli RNase HII .			3	1
ORF138	161527	162153	5'guanylate kinase (gmk)	U32848	Haemophilus influenzae	385	43
ORF139	162144	162443	putative			-	:
ORF140	162437	164098	methionyl-tRNA synthetase	AB004537	Schizosaccharomyces pombe	198	25
ORF141	165451	164228	exodeoxyribonuclease V (recD)	U32811	Haemophilus influenzae	432	32
ORF142	166349	165411	putative				
ORF143	166949	168442	putative		,		
ORF144	169416	171029	putative				
ORF145	170857	171459	putative			-	:
ORF146	172652	173428	putative biotin-protein ligase	Z97992	Schizosaccharomyces pombe	767	4
ORF147	174626	173439	putative		1		
ORF148	174816	175613	putative				
ORF149	175598	175954	putative				
ORF150	175958	176935	putative				

ORF	Begin	End	Homology	<b>a</b>	Species	Score	%I
ORF151	177708	176938	orf 3'of chaperonin homolog hypB [Chlamydia psittaci, pigeon strain P-1041, Peptide Partial, 98 aal	S40172	Chlamydia psittaci	376	74
ORF152	177128	177376	putative				
ORF153	179472	177841	putative	M69217	Chlamydia pneumoniae	2678	3 8
ORF154	179822	179517	putative	M69217	Chlamydia pneumoniae	498	3 3
ORF155	181793	179943	Pz-peptidase	D88209	Bacillus licheniformis	1088	<u></u>
ORF156	182628	181876	o247; This 247 as orf is 51 pct identical (0	AE000174	Escherichia coli	401	42
			gaps) to 117 residues of an approx. 160 aa				
ORF157	184420	183074	glutamate-1-semialdehyde 2,1-	X53696	Escherichia coli	823	41
			aminomutase			1	1
ORF158	184988	184467	ORF 0211	U28377	Escherichia coli	<u> </u>	47
ORF159	185483	185112	hypothetical protein	D90906	Synechocystis sp.	16	33
ORF160	185902	185483	ribose 5-phosphate isomerase	U28377	Escherichia coli	Ξ	4
ORF161	186174	185839	ribose 5-phosphate isomerase A	U32729	Haemophilus influenzae	190	46
			(SP:P27252)		-		,
ORF162	187720	186587	hypothetical	D83026	Bacillus subtilis	536	47.7
ORF163	188318	190933	ATP-dependent protease binding subunit	M29364	Escherichia coli	2010	23
ORF164	191090	191635	putative		=		
ORF165	191547	192743	putative				
ORF166	192969	193469	putative				
ORF167	194044	193610	putative			250	5
ORF168	194196	195809	unknown	Z84395	Mycobacterium tuberculosis	747	75
ORF169	196088	198073	DNA ligase (EC 6.5.1.2)	M24278	Escherichia coli	131/	40
ORF170	198132	199454	putative				
ORF171	199351	202818	putative		-	6	7
ORF172	204552	202999	PcpB	U60175	Sphingomonas chlorophenolica	08	4
ORF173	205648	204692	putative			1,000	:
ORF174	205807	207327	leucine tRNA synthetase	AF008220	Bacillus subtilis	35	ر ا
ORF175	207182	207775	leucyl-tRNA synthetase	X06331	Escherichia coli	363	7
ORF176	207779	208267	transfer RNA-Leu synthetase	M88581	Bacillus subtilis	782	5
ORF177	208267	209577	KDO transferase	Z31593	Chlamydia pneumoniae	7077	3

ORF	Begin	End	Homology	8	Species	Score	%[
ORE178	211807	211271	KDO-transferase	X80061	Chlamydia psittaci	105	38
ORF179	212188	211844	putative				
ORF180	214079	212448	pyrophosphate-dependent	Z32850	Ricinus communis	1003	45
			phosphofructokinase beta subunit	0007711		-	-
ORF181	214907	214083	CinI	U44893	Butyrivibrio fibrisolvens	=	<b></b>
ORF182	216154	215429	putative				
ORF183	216115	216678	putative				
ORF184	216728	217282	putative		9		
ORF185	217267	217866	putative				
ORF186	218593	218261	putative				
ORF187	219821	218994	putative				
ORF188	221382	220309	putative			;	97
ORF189	222719	221433	GMP synthetase	M10101	Escherichia coli		\$ 5
ORF190	223521	222724	IMP dehydrogenase	X66859	Acinetobacter calcoaceticus	8//	28
ORF191	224499	225008	putative				
ORF192	225140	225559	putative				
ORF193	225555	226802	putative				
ORF194	227800	226892	putative			1	
ORF195	228335	228072	putative				
ORF196	229251	228643	putative			1,000	1
ORF197	230983	229622	YqhX	D84432	Bacillus subtilis	1386	ဂ္ဂ
ORF198	231483	230983	acetyl-CoA carboxylase biotin carboxyl	U38804	Porphyra purpurea	961	75
			carrier protein	100174		200	33
ORF199	232063	231509	elongation factor P	D64001	Synechocystis sp.	707	75
ORF200	232739	232053	pentose-5-phosphate-3-epimerase	D90911	Synechocystis sp.	403	5
ORF201	233166	234356	putative				
ORF202	233518	233165	putative				
ORF203	234536	235186	ORF2	L35036	Chlamydia psittaci	570	3
ORF204	235379	236689	putative				
ORF205	236680	237618	putative			-	
ORF206	237521	238345	putative			1	
ORF207	238281	238973	putative				
ORF208	238871	240115	putative				

ORF	Begin	End	Homology	<b>a</b>	Species	Score	?
ORF209	240191	241564	putative				
ORF210	242281	241604	YaiZ	D84432	Bacillus subtilis	379	39
ORF211	242933	242274	gaps) to 208 residues of an approx. 232 aa	AE000284	Escherichia coli	382	45
00000	242416	77001	profiting repressor profein (arox)	132800	Haemophilus influenzae	229	46
ORF212	243410	244531	sialoalvoontotease	U15958	Pasteurella haemolytica	595	53
ORF213	242500	246021	oligopeptide permease homolog AII	AF000366	Borrelia burgdorferi	457	34
ORF215	246330	247811	OppAIV	AF000948	Borrelia burgdorferi	453	35
ORF216	247831	249174	OppA gene product	X56347	Bacillus subtilis	255	37
ORF217	249437	251038	dciAE	X56678	Bacillus subtilis	469	37
ORF218	251325	252212	OppB gene product	X56347	Bacillus subtilis	652	42
ORF219	253156	254007	oligopeptidepermease	X89237	Streptococcus pyogenes	574	48
ORF220	253974	254852	ATP binding protein	L18760	Lactococcus lactis	433	8
ORF221	255258	256094	KDO-transferase	X80061	Chlamydia psittaci	106	46
ORF222	256640	257455	putative			,	
ORF223	257502	258239	2-OXOGLUTARAT	A47930	Spinacia oleracea	636	77
ORF224	257869	257501	putative		÷		]:
ORF225	259248	260897	pyrophosphate-fructose 6-phosphate 1-	M55191	Solanum tuberosum	1055	4
			phosphotransferase beta-subunit		NA.		
ORF226	262753	261788	putative				
ORF227	263059	262757	putative				
ORF228	264375	263182	putative				
<b>ORF229</b>	265985	264747	putative				
ORF230	266637	266059	putative				
ORF231	267338	266538	putative				
ORF232	267922	267473	putative			30,	;
ORF233	269647	270771	tRNA guanine transglycosylase	L33777	Zymomonas mobilis	628	44:
ORF234	272777	273145	ORF 4	D00624	Bacteriophage chp1	IOO	41
ORF235	273253	273636	putative		Y		
ORF236	273705	273977	putative				
ORF237	276016	275717	putative				
<b>ORF238</b>	276439	276020	putative				

	End	Homology	A	Species	Score	%1
277253		putative				
277599 p	<u> </u>	putative				
	ā	putative			,	,
278554 F	ഥ	FbpC	U33937	Neisseria gonorrhoeae	312	ئر
	<u>a</u>	putative		1		
280849 p	_	putative		,	;;	5
282325 C	)	CMP-2-keto-3-deoxyoctulosonic acid	U15192	Chlainydia trachomatis	63/	<u>2</u>
S	S	synthetase			0000	7
284069 C		TP synthetase	U15192	Chlamydia trachomatis	7007	8
		ORF3	U15192	Chlamydia trachomatis	453	3
285775 g	20	glucose 6-phosphate dehydrogenase	U83195	Chlamydia trachomatis	1263	
285987 g	oc	glucose 6-phosphate dehydrogenase	U83195	Chlamydia trachomatis	519	6/
	50	glucose-6-phosphate dehydrogenase	D88189	Actinobacillus	216	04
i	· <u>··</u>	isozyme		actinomycetemcomitans		T
287002 pr	ď	putative				
287451 put	Ē	ıtative				
	ם	putative				
	Ē	putative				T
289213 pi	Ā	putative				
290254 pi	Ē	putative		4		1
	80 5	gamma-D-glutamyl-L-diamino acid	X64809	Bacillus sphaericus	95	39
292755 S	S	ScoS9	U43429	Streptomyces coelicolor	233	45
293272 r	-	ribosomal protein L13 (rpL13)	U32823	Haemophilus influenzae	364	47
293953 g	8	glutamine transport ATP-binding protein Q	U67524	Methanococcus jannaschii	387	46
	d	putative				
	a	putative			;	Ţ
	3	conserved hypothetical protein	AE000586	Helicobacter pylori	<u>\$</u>	46
297879 E		putative		4		
298253	$\Box$	putative		, i		
	$\dashv$	putative				
	-	putative				
300702	4	putative				

%I					 80 				42	42	42	ļ	9							44	39	31	48	59		]	43	37	4	62		33
Score					250				156	547	403	1	152							141	154	784	28	1838			376	356	545	1194		479
Species					Escherichia coli				Sinorhizobium meliloti	Bacillus subtilis	Bacillus firmus		Bacillus firmus							Schistosoma mansoni	Bacillus subtilis	Bacillus subtilis	Aeromonas salmonicida	Clostridium acetobutylicum			Helicobacter pylori	Escherichia coli	Haemophilus influenzae	Escherichia coli		Escherichia coli
A					AE000232				U81296	D84432	U61168		U61168							AF006678	X17014	D26185	L47978	U35453			AE000654	U70214	U32744	AE000299		AE000246
Homology	putative	putative	putative	putative	f311; This 311 as orf is 22 pct identical (13	gaps) to 186 residues of an approx. 488 aa	protein YACA BACSU SW: P37563; pyul	ofD21139	survival protein surE	YqfU	3-octaprenyl-4-hydroxybenzoate carboxy-	lyase	4-hydroxybenzoate octaprenyltransferase	putative	putative	putative	putative	putative	putative	lysophospholipase homolog	dnaZX	unknown	DNA gyrase	DNA gyrase subunit B	putative	putative	outer membrane protein	hypothetical	ATP-binding protein (abc)	f374; This 374 aa orf is 30 pct identical (9	gaps) to 102 residues of an approx. 512 aa	Xas A
End	301571	302437	302745	303852	305223				306236	307439	307458		308037	310180	311214	311253	311780	312772	313377	314665	314755	315531	316156	318676	321098	321710	322366	323181	323856	326410		327836
Begin	302440	302838	303335	304394	304606			-	305394	306501	308033		308924	309485	310426	311597	312772	313425	313646	313937	315576	316157	318657	321042	321445	322309	323190	323843	324878	325340		326433
ORF	ORF269	ORF270	ORF271	ORF272	ORF273				ORF274	ORF275	ORF276		ORF277	ORF278	ORF279	ORF280	ORF281	ORF282	ORF283	ORF284	ORF285	ORF286	ORF287	ORF288	ORF289	ORF290	ORF291	ORF292	ORF293	ORF294		ORF295

ORF	Begin	End	Homology		Species	Score	%I
ORF296	328465	327839	putative		•		П
ORF297	329360	328857	putative				
ORF298	330907	329357	putative				7
ORF299	332455	330956	MgtE	U18744	Bacillus firmus	203	္က
ORF300	334536	332395	putative				
ORF301	336091	334877	putative				T
ORF302	336103	337302	putative				
ORF303	338129	338830	putative				T
ORF304	338965	339501	putative		7		
ORF305	339508	340143	putative		1		
ORF306	340247	342967	putative				1
ORF307	343385	343810	cAMP-dependent protein kinase type I	U75932	Rattus norvegicus	102	37
			regulatory subunit				1
ORF308	344171	343935	acyl carrier protein (acpP)	AE000570	Helicobacter pylori	86	2
ORF309	345082	344330	3-ketoacyl-ACP reductase	U39441	Vibrio harveyi	298	84
ORF310	346005	345082	malonyl-CoA:Acyl carrier protein	U59433	Bacillus subtilis	538	45
			transacylase		7	62.0	T
ORF311	346784	346437	beta-ketoacyl-acyl carrier protein synthase	AE000540	Helicobacter pylori	2/3	? ?
0.0000	247020	246716	Lots Letonal poul carrier protein conthace	M77744	Escherichia coli	265	63
OKF-312	34/029	340/13	Dela-Ketoacyt-acyt carriet protein symmese				
ORF313	347034	347723	recombination protein	D90916	Synechocystis sp.	363	42
ORF314	348075	350459	putative				
ORF315	350598	351071	putative				
ORF316	351075	352175	rifampicin resistance protein	L22690	Rickettsia rickettsii	495	94
ORF317	353291	352230	putative			1	:
ORF318	353442	354467	pyruvate dehydrogenase E1 component,	D90915	Synechocystis sp.	571	<del>4</del>
OB F319	354451	354933	nymyate dehydrogenase E1 beta subunit	U09137	Arabidopsis thaliana	495	59
ORF320	355000	355449	pyruvate dehydrogenase El component,	U38804	Porphyra purpurea	336	47
OPE321	355448	356743	F23B12.5	Z77659	Caenorhabditis elegans	759	46
OB 52.1	355053	355642	nitative		- 21		
777.710	00000	4:000					1

ORF	Begin	End	Homology	αı	Species	Score	% %
		200774	1	300771	Homo conions	2193	57
OKF323	359310	350827	glycogen phospholytase D mitative	0.470	around a formation of the state		
JRF125	359575	359908	putative				
ORF326	361290	359947	DnaA	D89066	Staphylococcus aureus	375	46
ORF327	363785	361362	hypothetical	U32781	Haemophilus influenzae	394	4
328	364496	363888	putative				
ORF329	364832	365290	putative				:
ORF330	365304	365669	idp	M76470	Escherichia coli	160	45
ORF331	366599	365667	NADPH thioredoxin reductase	AC002329	Arabidopsis thaliana	975	8
ORF332	367291	369030	ribosomal protein S1 (rpS1)	U32801	Haemophilus influenzae	1209	4
ORF333	369134	369808	NusA	U74759	Chlamydia trachomatis	995	87
ORF334	369917	370438	NusA	U74759	Chlamydia trachomatis	760	87
ORF335	370365	372647		U74759	Chlamydia trachomatis	2173	61
ORF336	372557	373066	initiation factor IF2-beta (infB; gtg start	X00513	Escherichia coli	333	39
OPE337	373020	373442	ORF6 gene product	Z18631	Bacillus subtilis	192	34
OPE338	373467	374195	tRNA pseudouridine 55 synthase	D90917	Synechocystis sp.	358	47
ORF339	374176	375099	hypothetical 34.6 kD protein in rpsT-ileS	AE000113	Escherichia coli	395	39
			intergenic region			103	5
ORF340	375676	375083	hypothetical GTP-binding protein in pth 3' region	AE000219	Escherichia coli	207	3
OR F341	376173	375634	hypothetical	U32723	Haemophilus influenzae	480	59
ORF342	376564	377643	YscU	U08019	Yersinia enterocolitica	538	37
ORF343	377956	379773	lcrD gene product	1 <i>LLL</i> 121	Yersinia enterocolitica	1302	47
ORF344	379781	380425	putative				
ORF345	380281	381000	putative				
ORF346	381008	381460	putative		-2:	-	1
ORF347	381460	383037	4-alpha-glucanotransferase	L37874	Clostridium butyricum	302	38
ORF348	383257	383523	ribosomal protein L28 (rpL28)	U32776	Haemophilus influenzae	175	55
ORF349	383553	385304	hypothetical protein	D90901	Synechocystis sp.	565	38
ORF350	385397	386458	comE ORF1	D64002	Synechocystis sp.	187	2
ORF351	387242	386514	putative		===		
ORF352	388764	387013	putative				

ORF	Begin	End	Homology	<u>e</u>	Species	Score	 
ORE353	390120	390932	methylenetetrahydrofolate dehydrogenase	D64000	Synechocystis sp.	588	53
ORF354	390919	391818	to YOJL_ECOLI SW: P33944 (122 aa) and aa 152-351 are 100 pct identical to your ECOLI SW: P33944 (122 band about the part of the period of the part of	AE000310	Escherichia coli	186	39
ORF355	392379	391885	small protein	D90914	Synechocystis sp.	387	46
ORF356	392582	392986	putative				
ORF357	392776	393684	putative			000	,
ORF358	394151	394804	RecF protein	D90907	Synechocystis sp.	737	34
ORF359	394928	395308	putative			-	
ORF360	395259	395990	putative				,
ORF361	397815	395953	hypothetical	U32773	Haemophilus influenzae	195	9
ORF362	398850	397831	H. influenzae predicted coding region	U32763	Haemophilus influenzae	08 	96
Op 5373	400085	300000	nitative				
OR F364	401245	400073	YtgC	AF008220	Bacillus subtilis	244	30
ORF365	401474	401136	putative				
ORF366	402199	401423	unknown	U52850	Erysipelothrix rhusiopathiae	534	94
ORF367	403193	402186	putative		=		
ORF368	403650	404165	putative				1
ORF369	404343	405914	adenine nucleotide translocase	Z49227	Arabidopsis thaliana	1280	ટ્ર
ORF370	405984	407327	putative				
ORF371	407712	408806	putative				
ORF372	410439	409075	putative				
ORF373	411826	410954	putative				3
ORF374	412482	414302	lepA gene product	X91655	Bacillus subtilis	1827	59
ORF375	415402	414407	6-phosphogluconate dehydrogenase,	U32737	Haemophilus influenzae	687	51
ORF376	415848	415237	6-phosphogluconate dehydrogenase, 6PGD	S67873	Ceratitis capitata	695	Z
			[Ceratitis capitata=medflies, Peptide, 481				
ORF377	417131	415866	tyrosyl-tRNA synthetase (tyrS)	J01719	Escherichia coli	821	45
ORF378	417258	417566	putative				
, , , , , ,							

ORF	Begin	End	Homology	a	Species	Score	% %
				00000		464	41
ORF379	418326	417454	whiG-Stv gene product	60/89X	Sireptovernciilium griseocarneum	ţ P	F
ORF380	420057	418426	FLHA gene product	X63698	Bacillus subtilis	455	\$ (
ORF381	420448	420720	ferredoxin IV	M59855	Rhodobacter capsulatus	1/4	S
ORF382	420980	421552	putative				
ORF383	421556	422029	putative				
ORF384	422461	422925	putative				
ORF385	423562	424320	putative				
ORF386	424250	424591	putative				
ORF387	424830	426047	putative				
ORF388	426240	427397	putative			077	17
ORF389	428841	430703	GcpE	D90908	Synechocystis sp.	136	14
ORF390	430694	431446	НЭХ	U50134	Escherichia coli	130	ર
ORF391	431597	432100	putative				
ORF392	432165	432779	putative			476	13
ORF393	433272	432832	dihydrolipoamide succinyltransferase	U32839	Haemophilus influenzae	4 C	5
			(sucB)	00000	77	223	45
ORF394	433925	433227	dihydrolipoamide succinyltransferase	U32839	Haemophilus injiuenzae	326	}
200340	07777	133034	aluba-ketoolutarate debydrogenase	U41762	Rhodobacter capsulatus	1530	4
OKF395	430070	438357	oxygen-independent coproporphyrinogen	AE000628	Helicobacter pylori	442	42
OCC 200			III oxidase (hemN)				
ORF397	440317	438518	putative			_	
ORF398	440001	440345	putative			9,	۲
ORF399	441233	440517	ORF f286	U18997	Escherichia coli	8	€
ORF400	440719	441012	putative				
ORF401	442192	441230	putative				
ORF402	442888	442343	putative				
ORF403	442371	442961	putative		-	100	30
ORF404	443578	443003	[karp] gene products	M86605	Chlamydia trachomatis	2 5	0 5
ORF405	444500	443526	aminopeptidase	D17450	Mycoplasma salivarium	7/2	2
ORF406	444842	444528	putative	0000	3.4	122	77
ORF407	445009	444743	putative	L39923	Mycobacterium teprae	133	3

ORF	Begin	End	Homology	Œ	Species	Score	%I
ORF408	445718	445182	putative				:
ORF409	445807	447804	Sulp	U18908	Zea mays	1307	22
ORF410	448738	447803	putative				3
ORF411	449628	448618	RuvB protein	U38840	Thermotoga maritima	845	2
ORF412	450298	450867	deoxycytidine triphosphate deaminase (dcd)	AE000554	Helicobacter pylori	573	28
ORF413	450713	451207	putative				7
ORF414	451211	452452	hemolysin	D90914	Synechocystis sp.	227	5
ORF415	452448	453659	similar to [SwissProt Accession Number p37908]	D90888	Escherichia coli	96	33
ODEATE	454843	453775	NifS gene product	L34879	Anabaena azollae	533	38
ORF417	455608	454865	hypothetical protein	D90908	Synechocystis sp.	371	36
ORF418	456243	457007	putative				
ORF419	457016	457708	putative		-	-	];
ORF420	458368	457979	unknown	D26185	Bacillus subtilis	152	36
ORF421	459496	458372	mutY homolog	U63329	Homo sapiens	466	46
ORF422	459493	460194	hypothetical protein	D90914	Synechocystis sp.	86	38
ORF423	461446	460355	putative				
ORF424	462298	461450	putative			- 000	1
ORF425	462444	463349	enoyl-ACP reductase	Y13861	Nicotiana tabacum	1008	3
ORF426	464241	463342	putative				
ORF427	464574	465065	putative				
ORF428	465129	465611	putative				
ORF429	465571	466317	putative			7,70	7,
ORF430	466317	467093	H. pylori predicted coding region HP0152	AE000536	Helicobacter pylori	047	2
ORF431	466999	467502	putative		- =	, ,	٠
ORF432	469691	467715	unidentified transporter-ATP binding	Z82044	Bacillus subtilis	490	£ 8
ORF433	470691	469660	acetyl-CoA carboxylase subunit	AF008220	Bacillus subtilis	/81	77
ORF434	472010	470709	putative				
ORF435	471545	471799	putative				
ORF436	472359	472045	putative				1
ORF437	473523	472732	orfl	X75413	Escherichia coli	313	7 5
ORF438	474889	473441	murE gene product	Z15056	Bacillus subtilis	6/9	3
ORF439	477323	475365	penicillin-binding protein 2	X59630	Neisseria meningitidis	451	42

ORF	Begin	End	Homology	a	Species	Score	% 1
ORF440	478496	477597	hypothetical protein	D90906	Synechocystis sp.	534	52
ORF441	478722	479273	putative				
ORF442	479277	479705	putative			601	
ORF443	480050	481450	chromosomal replication initiator protein	D90909	Synechocystis sp.	56/	9
		0.000	Dilan	1135673	Rorrelia huradorferi	157	37
ORF444	481469	482053	Orth	0330/3	DOITEHU DUI KUOTIETT		
ORF445	482600	482025	putative		17.7	108	9
ORF446	482654	484204	NADH:ubiquinone oxidoreductase subunit B	23/111	Vibrio alginoiyiicus	100	7
ORF447	484211	485170	NADH:ubiquinone oxidoreductase	U32702	Haemophilus influenzae	258	8
	011101	405030	MADIfunianiana evidoreductase	737111	Vibrio alginolyticus	543	55
ORF448	485170	462636	INACTION IN CONTROL OF Na+-translocating	D49364	Vibrio alginolyticus	488	48
OKF 443	400013	20000	NADH-quinone reductase		,		
ORF450	486976	486638	putative				
ORF451	489071	487764	putative				
ORF452	489341	489090	putative				
ORF453	489958	489152	putative				
ORF454	490549	489962	putative				
ORF455	491163	490522	putative				T
ORF456	491396	491112	putative				
ORF457	492121	491390	putative			2270	77
ORF458	492304	494838	ClpC adenosine triphosphatase	U02604	Bacillus subtilis	0/57	040
ORF459	495943	494822	hypothetical protein in purB 5' region	AE000213	Escherichia coli	/76	2
ORF460	496011	496565	putative				
ORF461	496569	497228	putative				
ORF462	497358	497834	putative				
ORF463	497770	498327	putative				
ORF464	499209	499589	putative				
<b>ORF465</b>	499520	499792	putative			1016	36
ORF466	500774	504169	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	CIZI	3
ORF467	504139	504600	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	319	4 5
ORF468	504865	506877	putative 98 kDa outer membrane protein	0 / 2499	Chiamyala psillaci	226	75

1
putative 98 kDa outer membrane protein
putative 98 KDa outer memorane protein
POMP90A precursor
hypothetical
putative 98 kDa outer membrane protein
putative
putative 98 kDa outer membrane protein
POMP91B precursor
putative 98 kDa outer membrane protein
POMP91A
putative 98 kDa outer membrane protein
putative
putative 98 kDa outer membrane protein
putative
putative
putative 98 kDa outer membrane protein
putative outer membrane protein
putative
putative 98 kDa outer membrane protein
POMP91A

ORF	Begin	End	Homology	A	Species	Score	%
ORF502	547293	546994	putative			Š	5
ORF503	549699	550523	putative 98 kDa outer membrane protein	U72499	Chlainydia psittaci	96	75
ORF504	550490	551551	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	523	2
ORF505	551448	552623	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	139	9 9
ORF506	552652	555117	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	487	\$4
ORF507	555029	555493	putative				T
ORF508	558006	555673	putative				
ORF509	559694	558162	putative				
ORF510	558208	558573	putative		4		
ORF511	561692	559899	putative				
ORF512	561412	561708	putative		_		,
ORF513	563942	561777	1,4-alpha-glucan branching enzyme	X73903	Streptomyces coelicolor	1/43	45
ORF514	564969	563950	putative			,	18
ORF515	566204	564936	YqeV	D84432	Bacillus subtilis	639	28
ORF516	567717	566302	putative GTPase required for high	U00005	Escherichia coli	989	4 [
			frequency lysogenization by bacteriophage				
			lambda				
ORF517	568526	80229	putative				
ORF518	569467	568742	putative				
ORF519	571065	569431	putative				
ORF520	571828	571118	arginine-binding periplasmic protein 1	AE000188	Escherichia coli	197	<del>.</del>
			precursor				
ORF521	572202	573308	putative				T
ORF522	573146	575056	putative			1	1
ORF523	575023	575916	carboxysome formation protein	D90901	Synechocystis sp.	25/	3
ORF524	577891	576497	putative				
ORF525	578914	578204	putative		_		
ORF526	579924	578857	putative			- 1	Ţ
ORF527	580187	579858	protein kinase C inhibitor	D90906	Synechocystis sp.	097	4
<b>ORF528</b>	580017	580406	putative			72.	7,
ORF529	581086	580187	Yer156cp	018917	Saccharomyces cerevisiae	1/0	24
ORF530	581367	581828	putative				
ORF531	581678	582367	putative				7

Begin	End	Homology	<b>a</b>	Species	Score 1%
582361	583428	putative			
584690	583431	putative			
585237	584950	putative			
585626	586888	hypothetical protein	D64004	Synechocystis sp.	805
586846	587907	putative			
589049	588180	putative			
590500	589301	putative			
590755	592458	aminoacyl-tRNA synthetase	L25105	Chlamydia trachomatis	2125
592526	592903	has homology to putative heat shock	L25105	Chlamydia trachomatis	324
		acetohitylicum: ORFA; putative			
592836	593747	Possible negative regulator of CIRCE	US2216	Chlamydia trachomatis	096
		element; Homologs in B. subtilis and		·	
502747	\$04208	Clostridia spp. referred to as mcA of otta	M62819	Chlamydia trachomatis	199
594331	595947	DnaK protein homolog; 71,550 Da; putative	M69227	Chlainydia pneumoniae	2619
595905	596309	DnaK protein homolog; 71,550 Da; putative	M69227	Chlamydia pneumoniae	674
596514	597215	putative			
597184	597957	vacB gene product	U14003	Escherichia coli	306
597755	598612	ORF-2	D11024	Shigella flexneri	168
598602	599204	homologous to DNA glycosylases;	D83026	Bacillus subtilis	374
599373	599939	putative		13	
600903	602072	hemolysin	X73141	Serpulina hyodysenteriae	362
602240	602587	hypothetical protein	D90908	Synechocystis sp.	182
602637	603272	putative		· ·	
603142	604512	putative			
604627	605853	conserved hypothetical protein	AE000579	Helicobacter pylori	423
605790	606620	putative			
606571	607281	putative	L14679	Lactococcus lactis	384
609004	607355	putative			

ORF	Begin	End	Homology	Œ	Species	Score	<u>~</u>
ORFSS8	610906	609932	putative				
ORFS 59	611786	611004	diaminopimelate epimerase	D90917	Synechocystis sp.	207	55
ORF560	612333	611746	ATP-dependent Clp protease proteolytic	D90915	Synechocystis sp.	389	44
ODESS	613807	612341	serine hydroxymethyltransferase	D90903	Synechocystis sp.	606	52
ORF562	615179	616279	putative				
ORF563	616610	617383	putative			1	1
ORF564	961819	617810	ORF 0328	U18997	Escherichia coli	413	5
ORF565	620004	618826	branched chain alpha-keto acid	M97391	Bacillus subtilis	889	41
•			dehydrogenase E2				T
ORF566	619649	619918	putative			200	1,
ORF567	621265	620021	Hypothetical protein	Y14083	Bacillus subtilis	/7/	2
ORF568	622359	621265	hypothetical	U32691	Haemophilus influenzae	294	77
ORF569	623420	622560	rRNA methylase	D90913	Synechocystis sp.	244	38
ORF570	624297	623335	hypothetical protein (SP:P39587)	U67605	Methanococcus jannaschii	147	35
ORF571	624773	624174	riboflavin synthase alpha chain	AE000261	Escherichia coli	424	20
ORF572	625029	625484	ORF 168	D28752	Synechococcus sp.	323	43
ORF573	625488	625883	YteA	AF008220	Bacillus subtilis	172	35
ORF574	625892	626395	signalpeptidase II	X78084	Staphylococcus carnosus	204	38
ORF575	626444	627790	D-alanine permease (dagA)	U32770	Haemophilus influenzae	999	33
ORF576	627912	628607	putative		_		
ORF577	628774	629697	putative				:
ORF578	629660	631639	POMP91A	U65942	Chlamydia psittaci	579	4
ORF579	631725	633551	putative				:
ORF580	633520	636957	putative 98 kDa outer membrane protein	U72499	Chlamydia psittaci	266	4
ORF581	637232	638098	adhesion protein	D90903	Synechocystis sp.	267	88
ORF582	640648	639593	GTP-binding protein	D90901	Synechocystis sp.	759	3
ORF583	640979	640728	50S ribosomal protein L27	U38804	Porphyra purpurea	265	જ
ORF584	641327	641007	50S ribosomal subunit protein L21	U18997	Escherichia coli	210	4
ORF585	641687	642283	hypothetical protein	D90906	Synechocystis sp.	76	65
ORF586	643023	642286	assimilatory sulfite reductase	L26503	Saccharomyces cerevisiae	284	45
ORF587	643330	643076	putative			9,	5
ORF588	643704	643351	ribosomal protein S10 (rpS10)	U32761	Haemophilus influenzae	349	6

Begin	End	Homology	a	Species	Score	<u></u>
845628	929269	franslation elongation factor EF-G (fusA)	AE000625	Helicobacter pylori	1991	58
645783	645538	elongation factor G (AA 1-691)	X16278	Thermus aquaticus thermophilus	170	8
646269	645793	ribosomal protein S7	Z11567	Chlamydia trachomatis	730	88
646751	646314	ribosomal protein S12 (AA 1-123)	X52912	Cryptomonas phi	485	6)
647848	647045	putative				]
648393	650336	ORF of prc gene (alt.)	D00674	Escherichia coli	554	42
651016	650420	hypothetical sulfur-rich protein	U41759	Chlamydia psittaci	301	20
652956	651289	60kDa CrP	X53511	Chlamydia pneumoniae	2951	<u>8</u>
653395	653126	9kDa CrP	X53511	Chlamydia pneumoniae	502	8
655740	654193	glutamyl-tRNA synthetase homolog	U41759	Chlamydia psittaci	2259	82
656508	996559	early stage-specific transcription	L13598	Chlamydia psittaci	999	62
		experimentally demonstrated; early		<u></u>		
		upstream open reading frame (EUO)			3	:
658140	657022	unknown	U41759	Chlamydia psittaci	950	4
660216	658525	RecJ recombination protein	U41759	Chlamydia psittaci	807	23
663238	660248	protein-export membrane protein SecD	D64000	Synechocystis sp.	413	4
664461	663157	putative				
665735	664635	putative				1
666212	666994	hypothetical protein	D64006	Synechocystis sp.	538	28
866999	667921	o298; This 298 aa orf is 33 pct identical (24	AE000238	Escherichia coli	253	45
	<del></del>	gaps) to 248 residues of an approx. 256 aa				
		protein CDSA ECOLI SW: P06466			400	70
606199	895899	cytidylate kinase	AE000193	Escherichia coli	400	40
668502	669203	hypothetical protein	D90915	Synechocystis sp.	225	3
669154	670893	arginyl-tRNA-synthetase	D64006	Synechocystis sp.	1365	49
672226	670853	UDP-N-acetylglucosamine enolpyruvyl	U32788	Haemophilus influenzae	642	40
		transferase (murZ)				
671137	671424	putative				
672453	673001	putative				
673072	674721	putative				
674549	674262	putative			3	,
675518	674796	ORF246 gene product	X59551	Escherichia coli	220	43
676083	675499	putative				

ORF	Begin	End	Homology	QI	Species	Score	%1
ORF617	676630	190919	putative				,
ORF618	677016	009929	ORF3	D10279	Bacillus subtilis	361	63
ORF619	677647	677015	peptide release factor 2	X99401	Bacillus firmus	427	43
ORF620	066219	678259	unknown	Z49939	Saccharomyces cerevisiae	175	84
ORF621	679444	260089	unknown	D26185	Bacillus subtilis	263	38
ORF622	680097	268089	unknown	D64126	Bacillus subtilis	206	45
ORF623	681637	680849	putative				
ORF624	681409	682281	putative				
ORF625	682453	682821	putative				
ORF626	682763	683902	sensor protein	L39904	Myxococcus xanthus	190	84
ORF627	684616	696889	putative				
ORF628	685169	684534	putative				T
ORF629	986589	685117	putative			1	1
ORF630	686278	687288	NtrC/NifA-like protein regulator	U17902	Escherichia coli	820	5
ORF631	687483	688151	putative				
ORF632	688740	689501	putative				
ORF633	690242	689622	putative		1	- 3	1
ORF634	690470	691126	unknown	Z48008	Saccharomyces cerevisiae	380	46
ORF635	692600	691497	putative				
ORF636	692674	695064	phenylalanyl-tRNA synthetase beta-subunit	U32810	Haemophilus influenzae	593	45
ORF637	695049	696032	putative				
ORF638	697964	696585	OppC-like protein	D85103	Synechococcus sp.	371	37
ORF639	699803	698274	OppB gene product	X56347	Bacillus subtilis	197	용
ORF640	701926	882669	AppA	U20909	Bacillus subtilis	324	43
ORF641	703196	702567	putative				T
ORF642	704221	703208	putative			;	1
ORF643	704240	705289	ferrochelatase	X73417	Arabidopsis thaliana	266	42
ORF644	706070	705300	histidine periplasmic binding protein P29	U58045	Campylobacter jejuni	128	<u> </u>
ORF645	706841	706254	conserved hypothetical protein	AE000592	Helicobacter pylori	155	37
ORF646	707596	706811	putative			200	7
ORF647	999802	707677	ADP-glucose pyrophosphorylase	X55650	Solanum tuberosum	292	3
ORF648	709793	709119	pyrE-F gene product	X71842	Arabidopsis thaliana	400	4

ORF	Begin	End	Homology	<b>Q1</b>	Species	Score	%I
ORF649	711523	710132	transcription termination factor	J01673	Escherichia coli	1251	09
ORF650	712236	711523	putative				,
ORF651	714734	712125	DNA polymerase I	J04479	Streptococcus pneumoniae	1334	5
ORF652	715759	714761	protease IV	U67512	Methanococcus jannaschii	101	3
ORF653	717538	715886	adenine nucleotide translocase	Z49227	Arabidopsis thaliana	832	£)
ORF654	719113	720243	replicative DNA helicase	D26185	Bacillus subtilis	776	4
ORF655	720590	722422	homologous to E.coli gidA	X62540	Pseudomonas putida	1575	22
ORF656	722406	723056	putative				!
ORF657	723551	723120	nucleoside 5'-diphosphate	305207	Myxococcus xanthus	451	62
•			phosphotransferase (EC 2.7.4.6)			- 1	:
ORF658	724246	723626	Holliday junction DNA helicase (ruvA)	U32716	Haemophilus influenzae	293	5
ORF659	724754	724251	crossover junction endodeoxyribonuclease	U32717	Haemophilus influenzae	296	53
			(ruvC)				
ORF660	725868	724900	putative			1	
ORF661	727115	726270	putative				1
ORF662	728126	727119	glyceraldehyde-3-phosphate dehydrogenase	U83198	Chlamydia trachomatis	1340	2
ORF663	728594	728208	ribosomal protein L17	L33834	Chlamydia trachomatis	439	22
OR F664	729614	728604	RNA polymerase alpha-subunit	L33834	Chlamydia trachomatis	1356	8
ORF665	779778	729533	RNA polymerase alpha-subunit	L33834	Chlamydia trachomatis	273	82
ORF666	730149	729751	ribosomal protein S11	L33834	Chlamydia trachomatis	562	8
OR F667	730539	730174	ribosomal protein S13	L33834	Chlamydia trachomatis	54	68
ORF668	731983	730598	homolog	L25077	Chlamydia trachomatis	1956	8
OR F669	732427	731996	ribosomal protein CtrL15e	M80325	Chlamydia trachomatis	563	11
ORF670	732917	732423	ribosomal protein CtrS5e	M80325	Chlamydia trachomatis	702	<b>2</b>
ORF671	733598	733320	ribosomal protein L6	M60652	Chlamydia trachomatis	316	87
ORF672	733869	733492	ribosomal protein L6	M60652	Chlamydia trachomatis	469	77
ORF673	734298	733900	ribosomal protein CtrS8e	M80325	Chlamydia trachomatis	572	82
ORF674	734858	734319	ribosomal protein CtrL5e	M80325	Chlamydia trachomatis	730	8
ORF675	735195	734863	ribosomal protein CtrL24e	M80325	Chlamydia trachomatis	420	2
ORF676	735578	735342	ribosomal protein CtrL14e	M80325	Chlamydia trachomatis	270	95
ORF677	735861	735604	ribosomal protein S17e	M80325	Chlamydia trachomatis	322	11
ORF678	736492	736079	50S ribosomal protein L16	D90905	Synechocystis sp.	439	8

ORF	Begin	End	Homology	a	Species	Score	 %I
ORF679	737192	736524	ribosomal protein S3	D64071	Actinobacillus actinomycetemcomitans	612	28
ORF680	737555	737211	ribosomal protein L22	Z21677	Thermotoga maritima	228	48
ORF681	738688	737837	50S ribosomal subunit protein L2	U18997	Escherichia coli	692	62
ORF682	739048	738713	putative				
ORF683	739736	739065	ribosomal protein L4	X67014	Bacillus stearothermophilus	308	9
ORF684	740477	739773	ribosomal protein L3	Z46265	Thermus aquaticus thermophilus	463	8
ORF685	740659	740958	putative				T
ORF686	741722	740721	putative				
ORF687	742789	741827	methionyl-tRNA formyltransferase	D64001	Synechocystis sp.	511	48
ORF688	743618	742782	UDP-N-acetylglucosamine acyltransferase	L22690	Rickettsia rickettsii	542	<del>2</del>
ORF689	744092	743634	(3R)-hydroxymyristol acyl carrier protein	D90910	Synechocystis sp.	339	55
			dehydrase			1	Ţ:
ORF690	744604	744107	UDP-3-0-acyl N-acetylglcosamine	D90902	Synechocystis sp.	287	45
005601	744053	744498	TIDP-3-O-acvl-GlcNAc deacetylase	U67855	Pseudomonas aeruginosa	262	51
OP F697	746608	744986	anolinonrotein N-acyltransferase (cute)	U32716	Haemophilus influenzae	194	20
ORF693	747085	746621	low homology to P14 protein of	D78189	Bacillus subtilis	235	37
			Heamophilus influenzar and 14.2 kDa				
_			protein of Escherichia coli				7
ORF694	747974	747219	polymerase III	M22996	Bacillus subtilis	081	¥ :
ORF695	748594	748169	hypothetical protein	D90914	Synechocystis sp.	091	43
ORF696	749145	748573	putative			1	
ORF697	749652	749957	trxA	L39892	Chlamydia psittaci	393	7
ORF698	750446	749979	spoU	L39892	Chlamydia psittaci	559	27
ORF699	751219	750446	mip	L39892	Chlamydia psittaci	948	8
ORF700	753042	751291	aspartyl-tRNA synthetase	D90910	Synechocystis sp.	1347	47
ORF701	754309	753020	histidinetRNA ligase	Z17214	Streptococcus equisimilis	757	4
ORF702	755120	756175	hexosephosphate transport protein	M89480	Salmonella typhimurium	870	49
ORF703	756120	756485	hexosephosphate transport protein	M89479	Escherichia coli	321	45
ORF704	756499	760227	DNA polymerase III alpha-subunit (dnaE)	AE000646	Helicobacter pylori	1977	42
ORF705	761217	760297	putative				
ORF706	761297	761809	putative				

)	Fud	Homology	9	Species	Score
761782	762282	putative			
762260	762895	putative			
762867	763316	hypothetical protein	D90908	Synechocystis sp.	177
763780	763325	putative			
763861	765168	DD-carboxypeptidase	M85047	Bacillus subtilis	292
608994	765697	fmu and fmv protein	D90902	Synechocystis sp.	130
768051	766888	putative			
768566	768321	putative			
769342	768551	putative			
0532	769378	putative			
771451	770804	putative			
773058	771847	3-phosphoglycerate kinase	U83197	Chlamydia trachomatis	1540
773094	773456	putative			
774376	773093	putative phosphate permease	U84890	Mesembryanthemum crystallinum	870
775123	774380	putative			
775398	774916	putative			
775046	776077	sporulation protein	M57689	Bacillus subtilis	869
776070	777041	was dppE	U00039	Escherichia coli	565
777964	777536	orf288; translated orf similarity to SWISS-	Y10436	Coxiella burnetii	256
		PROT: YGIZ_PSEPU hypothetical 32.4			
778176	777904	B.subtilis genes rpmH, mpA, 50kd, gidA	X62539	Bacillus subtilis	112
		and gidB			
778621	779334	putative			
781173	780307	f406; This 406 aa orf is 28 pct identical (12	AE000263	Escherichia coli	603
		gaps) to 264 residues of an approx. 440 aa			
781526	781116	f406; This 406 as orf is 28 pct identical (12	AE000263	Escherichia coli	258
		gaps) to 264 residues of an approx. 440 aa			
782784	781555	f423; This 423 aa orf is 29 pct identical (1	AE000263	Escherichia coli	197
		gaps) to 172 residues of an approx. 488 aa			

ORF	Begin	End	Homology	ω	Species	Score	%I
			), 14(), 1, 1, 1, 1, 1, 1, 1, 1, 1, 1, 1, 1, 1,	1130004	Down him mismired	597	53
ORF731	783572	782805	hypothetical chloroplast OKF 10	D64004	Symechocystis sn	1720	62
ORF 732	785032	785381	ABC transporter submitt	200	.45		
OKF /33	700412	000001	nta transfer	Y14206	Streptomyces coelicolor	148	55
ORF /34	788944	788528	penicillin-binding protein 3	X84053	Pseudomonas aeruginosa	148	38
ORF736	789758	788901	putative				
ORF737	790332	791504	major outer membrane protein	M64064	Chlamydia pneumoniae	2028	66
ORF738	791846	792721	ribosomal protein S2	U60196	Chlamydia trachomatis	904	2
ORF739	792724	793569	elongation factor Ts	U60196	Chlamydia trachomatis	1023	7
ORF740	793580	794323	UMP kinase	U60196	Chlamydia trachomatis	168	72
OR F741	794304	794843	ribosome-releasing factor	U60196	Chlamydia trachomatis	673	73
ORF742	795217	795732	unknown	D26185	Bacillus subtilis	105	42
ORF743	795722	796795	unknown	D26185	Bacillus subtilis	208	33
ORF744	798735	797053	putative	L33796	Vibrio cholerae	386	34
ORF745	799823	798681	putative				
ORF746	799297	799578	putative			]	1
ORF747	801313	799808	Pkn5	U40656	Myxococcus xanthus	345	8
ORF748	802453	801332	putative				T
ORF749	803299	802457	putative				
ORF750	803811	803290	putative				1
ORF751	805151	803826	YscN	U02499	Yersinia enterocolitica	1185	2
ORF752	805860	805156	putative				
ORF753	806604	806332	putative				
ORF754	806913	809908	putative				
ORF755	808222	806903	putative				
ORF756	808751	808146	putative				
ORF757	809437	808673	putative				
ORF758	809939	809454	putative			,	5
ORF759	811235	810213	delta-aminolevulinate synthase (EC	M30785	Escherichia coli	7/1	€
			2.3.1.37)	277071	The state of the s	28.4	38
ORF760	811779	813056	DNA gyrase subunit B	U35453	Closifiaium acetobutyticum	100	ร
ORF761	812890	812516	putative			;	9
ORF762	812954	813583	DNA gyrase subunit B	8016IZ	Spiroplasma citri	3/1	2

Begin	End	Homology	8	Species	Score	%I
813587	815023	gyrA	X92503	Mycobacterium smegmatis	414	55
815420		putative				
816036	817010	orf-X; hypothetical protein; Method:	U48870	Bacillus subtilis	895	47
017111	917256	unchourn	7.74024	Mycobacterium tuberculosis	114	34
817701	818609	3-deoxy-d-manno-octulosonic acid 8-	Z50747	Chlamydia psittaci	1112	78
01//21	00010	nhosnbate synthetase	;			
818609	819094	protein of unknown function	Z50747	Chlamydia psittaci	545	65
819104		ATP binding protein	U72493	Chlamydia trachomatis	1099	88
820722		putative				
822313		putative				
823503		putative				
823678		putative				
825461	826312	putative				
827280		putative				
828604		76 kDa protein	L23921	Chlamydia pneumoniae	2179	2
830026		76 kDa protein	L23921	Chlamydia pneumoniae	1162	2
831047	-	mviB homolog	U50732	Chlamydia trachomatis	982	28
831725		mviB homolog	U50732	Chlamydia trachomatis	740	65
832220	-	T05H10.2	Z47812	Caenorhabditis elegans	407	34
833851		ribosomal protein S4 (rps4)	AE000633	Helicobacter pylori	372	53
834068		This ORF is homologous to a 40.0 kd	L22217	Mycoplasma-like organism	377	49
		hypothetical protein in the httB 3' region		<u> </u>		
835792	835127	uridine kinase	L31783	Mus musculus	436	43
837624	836116	ORF f397	U29581	Escherichia coli	92	38
838951	840882	putative				
840869		exodeoxyribonuclease V (recB)	U32811	Haemophilus influenzae	409	8
841989	-	DNA helicase II	U39703	Mycoplasma genitalium	110	46
843242		exodeoxyribonuclease V (recB)	U32811	Haemophilus influenzae	196	9
845018		MreC protein	M31792	Escherichia coli	76	2
846174	844990	aspartate aminotransferase (aspC)	X03629	Escherichia coli	754	9 2
848509		GreA	U02878	Rickeitsia prowazekii	190	35

Begin	End	Homology	8	Species	Score	%1
848568	849014	putative				į
849082	850488	NADH:ubiquinone oxidoreducatase subunit A (GP-737111 2)	U32702	Haemophilus influenzae	445	3/
851512	850574	porphobilinogen synthase	U38348	Chlorobium vibrioforme	692	45
852064	852447	putative				
852398	853690	putative			9	7
855118	854243	geranylgeranyl pyrophosphate synthase	D85029	Arabidopsis thaliana	408	41
855751	855128	f147; This 147 aa orf is 26 pct identical (1	AE000143	Escherichia coli	/81	30
		gaps) to 99 residues of an approx. 728 aa				
06/661	00000	membrane associated regulatory protein	M28368	Salmonella typhimurium	172	36
856551	629050	Illeliforation function	732530	Chlamydia trachomatis	842	35
856/30	626230	avodeovimihoniiclease V (recD)	U32811	Haemophilus influenzae	182	51
850501	860205	exonuclease V alpha subunit (AA 1-608)	X04582	Escherichia coli	235	45
861132	860284	nitative				
861426	861163	30S ribosomal protein S20	Z67753	Odontella sinensis	153	4
861701	862921	putative			.,,,,	3
863026	864798	major sigma factor	U04442	Chlainydia psittaci	7001	4
864831	865256	putative			1	٩
865226	866581	dihydropterin pyrophosphokinase	Y08611	Pisum sativum	455	δ <del>4</del>
		/dihydropteroate synthase		D	212	P
866562	867119	dehydrofolate reductase, type I (folA)	U32772	Haemophilus influenzae	202	25
867025	867816	M. jannaschii predicted coding region M10768	U67522	Methanococcus Jannaschii	/07	<b>ဂ</b>
867820	868497	putative				
869743	868661	RecA	U16739	Chlamydia trachomatis	1512	$\approx$
870633	870094	unknown function	Z32530	Chlamydia trachomatis	308	3
871929	870646	unknown function	Z32530	Chlamydia trachomatis	1410	8
872538	872086	putative		4.7		
873908	872517	putative				5
874281	874670	nifR3-like gene product	Z37984	Azospirillum brasilense	181	3/5
874582	875286	ORF1 gene product	X62399	Escherichia coli	30/	7 5
877857	875377	DNA topoisomerase I	L27797	Bacillus subtilis	1488	3

ORF	Begin	End	Homology	<b>a</b>	Species	Score	%
ORF820	878446	879255	putative			250	:
ORF821	880635	879268	sigma factor (ntrA) (AA 1-502)	X05888	Azotobacter vinelandii	/67	<b>1</b>
ORF822	882524	880593	DNA helicase II	D90906	Synechocystis sp.	1140	3 :
ORF823	882612	883319	ipa-57d gene product	X73124	Bacillus subtilis	100	7
ORF824	884155	883538	hypothetical protein	D90915	Synechocystis sp.	344	33
ORF825	884340	885611	19/20 residue stretch (32-51) identical to N-	L19954	Bacillus subtilis	456	37
			terminal putative signal sequence of				
			unknown, partly cloned B. subtilis gene.;				
		,	putative			,	
ORF826	885722	887302	heat shock protein	L12004	Chlamydia trachomatis	SIS.	<u>ئ</u> ر
ORF827	887587	888153	bas1 protein	Z34917	Hordeum vulgare	474	2
ORF828	888627	888220	putative				
ORF829	889330	888716	hypothetical protein	Y14079	Bacillus subtilis	223	5
ORF830	889898	889323	peptidoglycan-associated lipoprotein	X65796	Escherichia coli	222	S :
ORF831	891190	868688	TolB	U32470	Haemophilus influenzae	280	33
ORF832	891828	891247	putative			1	-
ORF833	892421	892017	exbD peptide	M28819	Escherichia coli	11	8
ORF834	893116	892421	inner membrane protein (tolQ)	U32722	Haemophilus influenzae	157	2
ORF835	892521	892925	putative				;
ORF836	893392	895419	inner membrane copper tolerance protein	Z36905	Escherichia coli	120	2
ORF837	895745	896527	unknown	D26185	Bacilius subtilis	381	41
ORF838	899968	897558	succinate dehydrogenase subunit C	Y08563	Paenibacillus macerans	253	<del>8</del>
ORF839	897565	899442	succinate dehydrogenase subunit A	Y08563	Paenibacillus macerans	1667	57
ORF840	899420	900229	succinate dehydrogenase subunit B	Y08563	Paenibacillus macerans	959	42
ORF841	903230	900237	putative				T
ORF842	905081	903234	putative			!	7
ORF843	906931	905045	sigma factor SibG regulation protein RsbU	D90905	Synechocystis sp.		સ
ORF844	907248	907832	putative				
ORF845	907784	908128	putative				
ORF846	908132	219806	putative				T
<b>ORF847</b>	685806	909320	putative				
<b>ORF848</b>	909405	911465	putative				
<b>ORF849</b>	911677	912360	putative				7

ORF	Begin	End	Homology	Œ	Species	Score	<u>%</u>
ORERSO	912303	912821	putative				
ORF851	912937	913983	putative				
ORF852	915128	914067	putative			7001	5
ORF853	916658	915303	enolase	L29475	Bacillus subtilis	1030	3
ORF854	915627	915376	enolase	U43738	Mycoplasma pneumoniae	077	3
ORF855	917707	916853	excinuclease ABC subunit B (uvrB)	U32804	Haemophilus influenzae	47/	<u>و</u>
ORF856	918837	917722	excinuclease ABC subunit B (uvrB)	U32804	Haemophilus influenzae	1029	4
ORF857	919868	918837	tryptophanyl-tRNA synthetase (trpS)	U32746	Haemophilus influenzae	3/6	₽
ORF858	920434	919880	putative				5
ORF859	921187	920438	ORF8	X82078	Chlamydia sp.	5 :	2
ORF860	921959	921195	hypothetical protein	X62475	Chlamydia psittaci	117	‡ ;
ORF861	923773	921995	Threonyl tRNA Synthetase	Z80360	Bacillus subtilis	14/0	4
ORF862	922146	922415	putative				
ORF863	923943	923674	putative				
ORF864	924077	925006	putative			T	
ORF865	925436	925083	putative				
ORF866	926524	925349	putative		Α,		
ORF867	927920	926433	putative				
ORF868	928319	927951	putative				
ORF869	928963	928334	putative			303	\$
ORF870	929248	930987	DNA mismatch repair protein (mutL)	U32692	Haemophilus influenzae	283	3 5
ORF871	930995	932059	YqhT	D84432	Bacillus sublilis	C#4	2
ORF872	932121	933515	putative				
ORF873	932881	932513	putative			150	22
ORF874	933485	935746	pulD (ttg start codon)	M32613	Klebsiella pneumoniae	017	ล่น
ORF875	935724	937082	epsE	M96172	Vibrio cholerae	200	ဂ္ဂ
ORF876	937229	938410	PilG	U32588	Neisseria gonorrhoeae	087	గ్గ
ORF877	938281	938805	putative				
ORF878	938809	939255	putative			+	
ORF879	939165	939782	putative		_		
ORF880	939760	940791	putative				
ORF881	940822	941106	putative				
ORF882	940977	941351	putative				

265 52 252 41 112 33			33 44 44 39 49 49	33 44 44 44 49 49 49 47	33 33 34 44 44 47 49 49 49	52 41 41 33 39 44 44 47 47 47 57	52 41 41 44 44 47 47 40 40 40 40 40 40 40 40 40 40 40 40 40	33 33 34 44 44 47 49 49 40 40 40 40 40 40 40 40 40 40 40 40 40	52 41 41 33 33 44 44 47 47 47 47 47 47 47 47 47 47 47	33 33 34 44 44 44 45 47 49 49 49 49 49 49 49 49 49 49 49 49 49		52 41 41 33 39 44 44 49 49 49 49 49 49 49 49 49 49 49	52 41 41 42 49 49 49 49 49 49 49 49 49 49 49 49 49	52 41 41 42 44 44 44 45 47 49 49 49 40 40 40 40 40 41 41 41 41 41 41 41 41 41 41 41 41 41
Aanthomonas campestras Yersinia enterocolitica	Naninomonas campestris Yersinia enterocolitica Erwinia amylovora	Kaninomonus campesiris Yersinia enterocolitica Erwinia amylovora Saccharomyces cerevisiae Bacillus subtilis	Naninomonus campesiris Yersinia enterocolitica Erwinia amylovora Saccharomyces cerevisiae Bacillus subtilis Staphylococcus aureus	a enterocolitica a amylovora romyces cerevisiae s subtilis lococcus aureus	Versinia enterocolitica  Erwinia amylovora  Saccharomyces cerevisiae  Bacillus subtilis  Bacillus cereus  Mycoplasma genitalium	Versinia enterocolitica Erwinia amylovora Saccharomyces cerevisiae Bacillus subtilis Mycoplasma genitalium Caenorhabditis elegans	Aaninomonus cumpesiris Yersinia enterocolitica Erwinia amylovora Saccharomyces cerevisiae Bacillus subtilis Mycoplasma genitalium Caenorhabditis elegans Lactobacillus delbrueckii	a enterocolitica a amylovora romyces cerevisiae ssubtilis lasma genitalium rhabditis elegans acillus delbrueckii us subtilis	a enterocolitica a amylovora a amylovora suromyces cerevisiae subtilis lococcus aureus lasma genitalium rhabditis elegans acillus delbrueckii subtilis subtilis subtilis	Versinia enterocolitica  Erwinia amylovora  Saccharomyces cerevisiae  Bacillus subtilis  Mycoplasma genitalium  Caenorhabditis elegans  Lactobacillus delbrueckii  Bacillus subtilis  Bacillus subtilis  Calamydia trachomatis	Erwinia amylovora  Erwinia amylovora  Erwinia amylovora  Bacillus subtilis  Mycoplasma genitalium  Caenorhabditis elegans  Bacillus subtilis  Bacillus subtilis  Caenorhabditis selegans  Caenorhabditis selegans  Caenorhabditis subtilis  Bacillus subtilis  Caulobacter crescentus  Bacillus subtilis	a enterocolitica a amylovora romyces cerevisiae ssubtilis lococcus aureus lasma genitalium rhabditis elegans acillus delbrueckii ssubtilis ydia trachomatis us subtilis ssubtilis ssubtilis	a enterocolitica a anylovora a anylovora romyces cerevisiae s subtilis lococcus aureus lasma genitalium rhabditis elegans ss subtilis	Erwinia amylovora  Erwinia amylovora  Bacillus subtilis  Staphylococcus aureus  Mycoplasma genitalium  Caenorhabditis elegans  Lactobacillus delbrueckii  Bacillus subtilis  Caulobacter crescentus  Bacillus subtilis
				2 222						2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2				
-	U56662	U56662 Z75104 M57435	U56662 Z75104 M57435 M73535	U56662 Z75104 M57435 M73535	U56662 Z75104 M57435 M73535 M73535 U398455	U56662 Z75104 M57435 M73535 M73535 X98455 Z68341	U56662 Z75104 M57435 M73535 M73535 U39680 Z68341 Z68341	275104 M57435 M57435 M73535 M73535 U39680 Z68341 Z68341	DS6662 Z75104 M57435 M73535 M73535 U39680 Z68341 Z68341 Z68341 Z68341 Z68341 Z68341 Z68341 Z78676 U11289 W72715	75104 275104 M57435 M73535 M73535 M73535 268341 268341 248676 U11289 X62539 U72715	275104 M57435 M57435 M73535 M73535 U39680 Z68341 Z68341 Z68341 U11289 X62539 U72715 U06928 L47648	Z75104 M57435 M57435 M73535 M73535 M73535 U39680 Z68341 Z48676 U11289 X62539 U72715 U06928 L47648	275104 M57435 M57435 M73535 M73535 U39680 Z68341 Z68341 Z68341 U11289 X62539 U72715 U06928 L47648 L47709 U18229	Z75104 M57435 M57435 M73535 M73535 U39680 Z68341 Z68341 Z68341 Z68341 U1289 V6128 U06928 U06928 U06928 U18229 U18229 U18229
		rogenase E3	rogenase E3	rogenase E3	rogenase E3	rogenase E3	rogenase E3 transferase E3	rogenase E3 transferase E3	rogenase E3 transferase E3 sid carrier K	rogenase E3 transferase E3 sid carrier K rboxylase	rogenase E3 transferase E3 cid carrier K k cboxylase cboxylase Escherichia coli	rogenase E3 transferase E3 id carrier K rboxylase troxylase Escherichia coli	rogenase E3 transferase E3 cid carrier K tboxylase tboxylase Escherichia coli 1; putative	rogenase E3 transferase E3 cid carrier k k tboxylase Escherichia coli a; putative
inve	utive utive J	putative putative HrcJ putative ORF YOR196c dihydrolipoamide dehydrogenase E3	putative Putative HrcJ putative ORF YOR 196c dihydrolipoamide dehydrogenase E3 subunit dihydrolipoamide acetyltransferase E3 subunit	ative J ative Tive Tive Autolipoamide dehydroger Adolipoamide acetyltransf unit unit Adolipoamide acetyltransf ative	utive J J strive Attive Attive F YOR 196c vdrolipoamide dehydroger unit ydrolipoamide acetyltransl unit taiive F F	utive J strive strive F YOR 196c vdrolipoamide dehydroger unit vdrolipoamide acetyltranst unit EF case G4.1	ative J Itive Since Sinc	putative  Putative  HrcJ  putative  ORF YOR 196c  dihydrolipoamide dehydrogenase J  subunit  dihydrolipoamide acetyltransferase subunit  putative  SNF  helicase  F01G4.1  putative  branched-chain amino acid carrier endonuclease III	HrcJ  HrcJ  Butative  Dutative  ORF YOR196c  Gihydrolipoamide dehydroger Subunit  dihydrolipoamide acetyltranst subunit  putative  FOIG4.1  putative  branched-chain amino acid ca endonuclease III homologous to E.coli 50K  honologous to E.coli 50K  honoshatidylserine decarboxy	putative Putative HrcJ Dutative ORF YOR196c dihydrolipoamide dehydrogenase subunit dihydrolipoamide acetyltransferas subunit putative SNF helicase F01G4.1 putative branched-chain amino acid carrier endonuclease III homologous to E.coli 50K phosphatidylserine decarboxylase putative	putative  HrcJ  putative  ORF YOR196c  dihydrolipoamide dehydrogenase E3 subunit  putative  SNF  helicase F01G4.1  putative  branched-chain amino acid carrier endonuclease III  homologous to E.coli 50K  phosphatidylserine decarboxylase putative  secretory component 28.2% of identity to the Escherichia coli	ative  ative  ative  ative  E YOR 196c  cydrolipoamide dehydrogenase E younit  ative  icase  1G4.1  ative  Inched-chain amino acid carrier achouclease III  mologous to E.coli 50K  sosphatidylserine decarboxylase ative  artive  Inched-chain amino acid carrier active  Inched-chain amino	ative ative ative ative FYOR196c ydrolipoamide dehydroger ydrolipoamide acetyltransf ydrolipoamide acetyltransf ydrolipoamide acetyltransf ative FF ative IG4.1 ative Inched-chain amino acid ca nched-chain amino acid ca ative Inched-chain amino acid ca ative refery Sphatidylserine decarboxy ative refory component 2% of identity to the Esch P-binding protein Era; put Iy(A) polymerase X-like protein	putative putative HrcJ putative HrcJ putative ORF YOR196c dihydrolipoamide dehydrogenase E3 subunit dihydrolipoamide acetyltransferase E3 subunit putative SNF helicase F01G4.1 putative branched-chain amino acid carrier endonuclease III homologous to E.coli 50K phosphatidylserine decarboxylase putative secretory component 28.2% of identity to the Escherichia coli GTP-binding protein Era; putative poly(A) polymerase CIP-binding protein Era; putative poly(A) polymerase CIP-Like protein ATP-dependent protease ATPase subunit
994 putative														
943994 944556 945389	945751	945751 948081 948915 949868	945751 948081 948915 949868 950959	945751 948081 948915 949868 950959	945751 948081 949868 949868 950959 952134 952165									
944413 945395 945853	946392	946392 947410 949871 951058	946392 947410 949871 951058	946392 947410 949871 951058 951249 95164	946392 947410 949871 951058 951249 951664 952674 953491	946392 947410 949871 951058 951249 951664 952674 953491	946392 947410 949871 951058 951249 951664 952674 953491 953324 955823	946392 947410 949871 951058 951249 951664 952674 953491 953491 955823 957082	946392 947410 949871 951058 951249 951664 952674 953491 955324 955324 955823 957082 957082	946392 947410 949871 951058 951249 952674 953491 953324 955324 955823 957082 957902 959376	946392 947410 949871 951058 951249 952674 953491 953491 955324 955324 957082 957082 957902 959376 960266 960855	946392 947410 949871 951058 951249 952674 953491 953324 955324 955324 955324 959376 960266 960855	946392 947410 949871 951058 951249 953491 953491 95324 955324 955324 955324 955324 959231 959231 959231 959231 959231 959231 959231 959231 959231	946392 947410 949871 951058 951249 952674 952674 953491 957082
ORF888 ORF889 ORF890	ORF891	ORF891 ORF892 ORF893 ORF894	ORF891 ORF892 ORF893 ORF894 ORF895	ORF891 ORF892 ORF893 ORF894 ORF895 ORF896	ORF891 ORF892 ORF893 ORF894 ORF895 ORF895 ORF896	ORF891 ORF892 ORF893 ORF895 ORF895 ORF896 ORF896 ORF899	ORF891  ORF892  ORF893  ORF894  ORF895  ORF896  ORF899  ORF899  ORF899	ORF891 ORF892 ORF893 ORF894 ORF895 ORF896 ORF896 ORF899 ORF900 ORF901	ORF891  ORF892  ORF893  ORF894  ORF896  ORF896  ORF899  ORF900  ORF901  ORF901	ORF891  ORF892  ORF893  ORF894  ORF896  ORF899  ORF900  ORF901  ORF901  ORF903  ORF903  ORF904  ORF905	ORF891 ORF892 ORF893 ORF894 ORF896 ORF896 ORF899 ORF900 ORF901 ORF903 ORF903 ORF904 ORF905 ORF906 ORF906	ORF891  ORF892  ORF893  ORF894  ORF895  ORF896  ORF800  ORF901  ORF901  ORF902  ORF904  ORF905  ORF905  ORF906  ORF906	ORF891  ORF892  ORF893  ORF894  ORF896  ORF899  ORF900  ORF901  ORF902  ORF904  ORF905  ORF906  ORF906  ORF906	ORF891  ORF893  ORF894  ORF894  ORF895  ORF896  ORF900

ORF	Begin	End	Homology	ΩI	Species	Score	<u>%</u>
						,	-
ORF912	970118	969762	ATP-dependent clp protease proteolytic component (clpP)	AE000591	Helicobacter pylori	362	3
ORF913	970593	970300	putative				
ORF914	971261	970542	putative				
ORF915	971680	971123	putative		1,		
ORF916	971876	975100	SNF	X98455	Bacillus cereus	778	49
ORF917	975419	916516	MreB protein	M96343	Bacillus subtilis	096	55
ORF918	976584	978320	phospho enol pyruvate carboxykinase	S56812.	Chlorobium limicola	1667	2
ORF919	089246	977231	putative				
ORF920	978399	980738	putative		27		
ORF921	980756	981928	putative				
ORF922	982974	981931	precursor protein (AA -22 to 371)	X52557	Chlamydia trachomatis	97	20
ORF923	984120	983119	NAD+ dependent glycerol-3-phosphate	L47648	Bacillus subtilis	618	43
			dehydrogenase				
ORF924	985502	984120	AgX-1 antigen [human, infertile patient, testis. Pentide 505 aal	S73498	Homo sapiens	254	34
ORF925	987180	985882	ORF 4	M72718	Bacillus subtilis	269	38
ORF926	987172	987444	putative				
ORF927	989846	989049	nifU-like protein	AE000542	Helicobacter pylori	302	31
ORF928	991048	989846	putative				
ORF929	991638	990955	phosphoglyceromutase	L09651	Zymomonas mobilis	471	53
ORF930	991794	992498	ORFX13	L09228	Bacillus subtilis	403	39
ORF931	993619	993041	biotin [acetyl-CoA-carboxylase] ligase	L47709	Bacillus subtilis	136	38
ORF932	993530	994792	rod-shape-determining protein	M22857	Escherichia coli	312	4
ORF933	995970	994795	cadmium-transporting ATPase	D64005	Synechocystis sp.	358	47
ORF934	996857	995739	ATPase	L28104	Transposon Tn5422	449	39
ORF935	997603	996782	putative				
ORF936	696866	997572	seryl-tma synthetase	Y09924	Staphylococcus aureus	851	42
ORF937	968866	1000023	orf2, homologue to B.subtilis ribG	X64395	Escherichia coli	965	9
ORF938	1000087	1001340	GTP cyclohydrolase II	D90912	Synechocystis sp.	1078	22
ORF939	1001357	1001818	riboflavin synthase beta subunit	U27202	Actinobacillus pleuropneumoniae	278	36
ORF940	1003288	1001873	putative				
ORF941	1003487	1004146	putative				

ORF	Begin	End	Homology	Œ	Species	Score	 %
OP F942	1004485	1005639	D-alanine glycine permease (dagA)	AE000603	Helicobacter pylori	394	33
OR F943	1005643	1005972	hypothetical protein MTCY180.08	Z97193	Mycobacterium tuberculosis	274	88
ORF944	1006784	1006116	similar to trithorax protein in final three	S18810	Caenorhabditis elegans	155	46
			exons	270102	D = ::     L +:   !:	406	38
<b>ORF945</b>	1007563	1006769	yycJ	D/8193	Bacillus sublitis	3 5	3
ORF946	1009226	1007568	YtpT	AF008220	Bacillus subtilis	766	4/
ORF947	1009989	1009336	putative				
ORF948	1015852	1016337	putative				
ORF949	1016561	1016181	putative				Ī
ORF950	1016297	1017532	putative				T
ORF951	1016802	1016452	putative			1	!
ORF952	1018993	1017701	phenolhydroxylase component	U32702	Haemophilus influenzae	909	4
ORF953	1019454	1019137	ORF	M63939	Escherichia coli	8	5
OR F954	1020764	1019562	pCTHom1 gene product	M94254	Chlamydia trachomatis	1185	65
ORF955	1021405	1021037	histone H1-like protein	M80324	Chlamydia psittaci	319	62
ORF956	1021821	1024286	phosphoprotein	L25078	Chlamydia trachomatis	739	4
ORF957	1024697	1024248	putative				
ORF958	1025569	1024508	protoporphyrinogen oxidase	U25114	Mus musculus	98	38
ORF959	1026969	1025590	oxygen independent coprophorphyrinogen	D90912	Synechocystis sp.	880	42
OBEGEO	1027789	1026947	uropomyvrinogen decarboxylase	M97208	Bacillus subtilis	372	38
ORF961	1031199	1027945	transcription-repair coupling factor (trcF)	U32805	Haemophilus influenzae	1584	42
ORF962	1031717	1031172	alanyi-tRNA synthetase	X95571	Thiobacillus ferrooxidans	76	31
ORF963	1033057	1031612	alanyl-tRNA synthetase	AE000353	Escherichia coli	688	6
ORF964	1033425	1033039	alanyl-tRNA synthetase (alaS)	AE000629	Helicobacter pylori	327	2
ORF965	1033784	1033200	alanyl-tRNA synthetase	X59956	Rhizobium leguminosarum	416	47
ORF966	1033963	1036038	transketolase	Z73234	Bacillus subtilis	1398	44
ORF967	1036945	1036010	AMP nucleosidase	AE000290	Escherichia coli	265	42
ORF968	1037110	1037679	elongation factor P	U14003	Escherichia coli	458	21
ORF969	1037696	1037944	putative				
ORF970	1038916	1037975	putative				;
ORF971	1040582	1039026	HSP60 chaperonin	X62914	Clostridium perfringens	784	21

ORF	Begin	End	Homology	<b>a</b>	Species	Score	~ %I
ORF972	1040997	1042337	PROBABLE UDP-N-ACETYLMURAMOYLALANYL-D-GLUTAMYL-2, 6-DIAMINOLIGASE (EC	AB001488	Bacillus subtilis	446	39
Op 5073	1042357	1043403	6.3.2.15). ORF-Y (AA 1-360)	X51584	Escherichia coli	582	45
ORF974	1043367	1044623	UDP-N-acetylmuramoylalanine-D-	U32793	Haemophilus influenzae	348	42
2000	1044207	1045362	glutamate ligase (murD)	Y14079	Bacillus subtilis	115	38
ORF975	1044607	1046538	spoVE gene product (AA 1-366)	X51419	Bacillus subtilis	479	35
OR F977	1046447	1047517	mur	Y13922	Enterococcus hirae	256	45
ORF978	1047521	1049956	UDP-N-acetylmuramate-alanine ligase	U32794	Haemophilus influenzae	756	38
02020	1050511	1050036		Z74024	Mycobacterium tuberculosis	78	4
ORF979	1050925	1050566	cycY gene product	U14003	Escherichia coli	179	34
OR F981	1051728	1051090	putative				7
OR F987	1051743	1052063	hypothetical protein	D90908	Synechocystis sp.	135	33
ORF983	1052101	1053126	trna delta(2)-isopentenylpyrophosphate	Z98209	Mycobacterium tuberculosis	441	37
70000	1054201	1052107	transferase	AE000579	Helicobacter pylori	826	44
OKF984	1054201	1055555	nitative				
ORFORK	1055483	1055908	putative		in .		
ORF987	1056609	1056965	YqeL	D84432	Bacillus subtilis	202	28
ORF988	1056961	1058232	beta-ketoacyl-ACP synthase	L13242	Ricinus communis	1200	ဂ္ဂ
ORF989	1058238	1058687	diadenosine tetraphosphatase	U30313	Homo sapiens	771	74
ORF990	1059371	1058727	inorganic pyrophosphatase (ppa)	AE000576	Helicobacter pylori	507	3
ORF991	1059526	1060578	leucine dehydrogenase LeuDH	U51099	Bacillus cereus	080	5
ORF992	1061553	1060579	3'(2'),5'-bisphosphate nucleotidase	U40433	Arabidopsis thaliana	335	24
ORF993	1061674	1062411	putative				1
ORF994	1062377	1064077	2-acylglycerophosphoethanolamine acyl	U29581	Escherichia coli	383	44
ORF995	1064116	1065243	7-keto-8-aminopelargonic acid synthetase	M29291	Bacillus sphaericus	200	35
200200	1067451	1065178	nri A	Y10304	Bacillus subtilis	1009	43
ORF996	106/451	10001/0	hin hin	23300			

国	End	Homology	a	Species	Score	<u>%</u>
1067376 putative	putati	ve				
1068706 putative	putativ	9/			1111	7
1068819 unknown	unknov	/m	U41759	Chlamydia psittaci	16	416
1070033 unknown	unknow	u	U41759	Chlainydia psittaci	381	2
-	unknow	1	U41759	Chlamydia psittaci	4C7	2
	putative		200000	G Of boarefie can	1007	48
1075864 lysyl-tk	lysyl-tk	lysyl-tKNA synthetase	1 14580	Bacillus subtilis	395	22
+	cvs-tRN	cvs-tRNA synthetase (cvsS)	U32693	Haemophilus influenzae	431	56
	putative				S. F.	1
1078672 ribonucle	ribonucle	ribonuclease P protein component (gtg start	M11056	Escherichia coli	0	0
1070944 30S riboso	30S riboso	30S ribosomal subunit protein \$14	U18997	Escherichia coli	260	50
	F18C12.2		Z75536	Caenorhabditis elegans	118	38
nd	putative				100	];
1081350 deoxyribodi	deoxyribodi	deoxyribodipyrimidine photolyase	J03294	Bacillus subtilis	687	44
1083235 DNA mism	DNA mism	DNA mismatch repair protein	U71154	Aquifex pyrophilus	735	48
1084632 DNA mism	DNA mism	DNA mismatch repair protein	D90909	Synechocystis sp.	202	3
	DNA prim	ase (dnaG)	U32735	Haemophilus influenzae	200	27 2
$\dashv$	DnaG		783860	Mycobacterium tubercutosis	777	2
$\dagger$	pulative					
1089800 putative 1089890 glycyl-tRN	putative glycyl-tRN	purative glycyl-tRNA synthetase	U20547	Chlamydia trachomatis	2569	48
1092889 putative	putative					
	phosphati	phosphatidylglycerophosphate synthase	U87792	Bacillus subtilis	163	3
1094192 glycogen	glycogen	glycogen (starch) synthase	D90899	Synechocystis sp.	5/4	€ 2
_	partial ctc	partial ctc gene product (AA 1-186)	X16518	Bacillus subtilis	98	3
	peptidyl-t	RNA hydrolase	U31570	Chlamydia trachomatis	378	53
-	ribosomal	ribosomal protein S6 (rps6)	AE000630	Helicobacter pylori	179	33
1097867 ribosoma	ribosoma	ribosomal protein S18 homolog; putative	M62820	Chlamydia trachomatis	324	2 8
1098392 putative	putative	putative heat shock protein ORF; putative	M62820	Chlamydia trachomatis	<u> </u>	2
1099279 putative	putative					
1101053 putative	putative					

Homology
1101107 putative
102116 glycerol-3-phosphate acyltransferase
O
1107249 putative
1108101 PlsX
1113370 putative 98 kDa outer membrane protein
1115071 lipid A disac
1116894 poly(A) polymerase
1120030 putative
-
1121430 glutamine amidotransferase; glucosamine
fructose-6-pk
1121866 L-glutamine:D-fructose-6-P
1122899 tyrosine-specific transport protein
1125564 putative
1127676 succinyl-CoA synthetase beta-subunit
1131336 putative
1132553 putative
-
1140102 Ydr430cp; CAI: 0.15

Begin	End	Homology	a	Species	Score	%
1141365	1140112	hypothetical 54.7 kD protein in udp 3'	AE000459	Escherichia coli	222	34
1142150	1141356	phosphatidylserine synthase (pssA)	AE000614	Helicobacter pylori	307	41
1142520	1145660	ribonucleotide reductase subunit M1	K02927	Mus musculus	1433	45
1145627	1146721	ribonucleoside diphosphate reductase, beta	AE000553	Helicobacter pylori	443	32
1146067	1147545	subunit (nrdb)	795398	Mycobacterium leprae	191	35
1147666	1148190	YtaB	AF008220	Bacillus subtilis	262	44
1148514	1148224	ORF2	U01958	Bacillus licheniformis	135	54
1149136	1148348	ORF2	M31827	Bacillus subtilis	268	9
1149702	1149166	putative				
1150031	1150591	unknown	Z85982	Mycobacterium tuberculosis	445	64
1150785	1151147	ribosomal protein L20 (AA 1-119)	X16188	Bacillus stearothermophilus	273	4
1151165	1152181	phenylalany-tRNA synthetase beta subunit	Z75208	Bacillus subtilis	777	9
1152522	1154591	putative				
1155666	1154566	putative		× 2		
1156743	1155670	putative		2		
1156859	1157815	hypothetical	U32723	Haemophilus influenzae	252	47
1157982	1160735	ATP-binding protein	U01376	Escherichia coli	1314	28
1162620	1160917	polynucleotide phosphorylase	AF010578	Pisum sativum	1416	22
1162970	1162590	polyribonucleotide phophorylase	U52048	Spinacia oleracea	312	53
1163532	1164020	orf150 gene product	X95938	Porphyromonas gingivalis	335	43
1163995	1164294	putative				
1165569	1165030	putative				
1166108	1165566	putative				
1166644	1166141	putative				
1167055	1168374	putative				;
1169218	1168337	methionine aminopeptidase	D64003	Synechocystis sp.	488	24
1169823	1169218	ORF 0197	U18997	Escherichia coli	281	3
1171324	1170572	putative		\$		
1172085	1171177	hypothetical	U32720	Haemophilus influenzae	79[	4 1
1172394	1173773	fumarase	D64000	Synechocystis sp.	1292	2
1175209	1173881	prs-associated putative membrane protein	U02424	Escherichia coli	270	3

ORF1087 11  ORF1088 11  ORF1090 11  ORF1092 11  ORF1093 11  ORF1094 11  ORF1095 11	1175555 1175778 1177177 1179156 1180045 1181942 11812296 1183844 1183844	1175127	L. mothatical protein in nth-nrs intergenic	AE000219	1. 1. 1.		1
	75778 77177 79156 80045 81942 82296 83844 84420	1177043	nypotitetical protein in pur-pra intergenie		Escherichia coli	278	46
	77177 79156 80045 81942 82296 83844 84420		hypothetical protein	Z96072	Mycobacterium tuberculosis	109	43
	79156 80045 81942 82296 83844 84420	1179048	putative				
	80045 81942 82296 83844 84420	1180085	penicillin tolerance protein (lytB)	U32781	Haemophilus influenzae	731	χ 24
	81942 82296 83844 84420	1180779	putative				
	82296 83844 84420	1180788	putative				
	83844 84420	1181961	putative				
	84420	1182300	putative				
-		1183848	putative				
_	1185382	1184366	putative				
ORF1097 11	1185858	1185226	putative				
-	1186164	1186481	putative				1
	1187386	1186484	site-specific recombinase	U92524	Salmonella typhimurium	401	48
	1187370	1189028	phophoglucoisomerase-like protein	L40822	Chlamydia trachomatis	1154	63
-	1189321	1190889	putative			100	,
_	1191142	1192146	NADP-malate dehydrogenase	L40958	Flaveria bidentis	775	46
	1191974	1191729	putative				
ORF1104 11	193815	1192991	putative		Y.		_]:
ORF1105 11	1195702	1194248	o460; This 460 aa orf is 46 pct identical (26	AE000256	Escherichia coli	1022	4
			gaps) to 458 residues of an approx. 488 aa				
			protein ARCD PSEAE SW: P18275				
ORF1106 11	1196303	1195716	putative				
ORF1107 11	1196831	1196337	putative				
	1197807	1196746	putative				
	1198740	1197883	putative			-	
-	1200232	1198721	shikimate 5-dehydrogenase	U67551	Methanococcus jannaschii	245	2
	1201286	1200135	3-dehydroquinate synthase (aroB)	U32705	Haemophilus influenzae	478	\$
_	1202386	1201259	2,3-dihydroxybenzoic acid	L29562	Vibrio anguillarum	780	2
-	1202901	1202350	putative				
ORF1114 12	1204162	1202816	5-enolpyruvylshikimate 3-phosphate	N67500	Methanococcus jannaschii	220	9
OPE1115 12	1203177	1203464	putative		-2		

1205028   1206392   1206392   1206392   1206392   1207872   12107852   1210703   1211870   1212742   12120630   1210710   12120630   1220630   1220630   1220630   1220630   1220630   1220630   1220972   1229972   1			operies		2
1206392	80 putative			1	Ţ;
1206742		A02587	unidentified	834	84
1207872	-	U32830	Haemophilus influenzae	243	37
1208852	-	U51868	Bacillus subtilis	109	41
1210518 1210703 1211870 1211870 1211870 1211870 1211870 1216493 1216493 1220068 1220068 1220068 1220068 1221645 122164 122164 122164 122164 122164 122164 122164 122164 122164 12216 12216 12216 12216 12216 1		U24147	Arabidopsis thaliana	892	52
1210703   1211870   1211870   1211870   12124046   1215551   1216493   1210068   1220068   1220068   1220069   1220000   1221645   12200000   12200000   12200000   12200000   12200000   122000000   12200000   12200000   12200000   12200000   12200000   12200000   12200000   12200000   12200000   122000000   122000000   122000000   122000000   122000000   122000000   122000000   122000000   122000000   122000000   122000000   122000000   122000000000   1220000000   1220000000000		U26428	Gallus gallus	237	34
1211870 1212742 1212742 1215551 1216493 1216493 1216493 1220068 1220068 1220068 1220068 1220680 1221645 1221645 1221645 1221645 1225900 1226528 1226528 1226528 1226972 1226972 1226972 1226972 1226972 1226972 1226972 1226972 1226972 1226972 1226972 1226972		U47017	Pseudomonas syringae pv. tabaci	345	37
1212742 1214046 1215551 1216493 1216493 1216493 1217183 1220068 1220068 12200630 1220630 1221645 1223894 1225000 1225000 1225000 1225972 1229972 47569 47569 53356 53356 53356 53356 53356		U67476	Methanococcus jannaschii	444	43
1214046 1215551 1216493 1216493 1220068 1220068 1220068 1220630 1220630 1221645 1223894 1223894 1225000 1225000 1225897 1229972 47569 49980 53356 53356 53356 53356 53356 53356		000000	Escherichia coli	473	4
1215551 1216493 121068 1220068 1220068 1220630 1221645 1221645 1221645 1221645 1221645 1222894 1225000 1226528 1226997 1226990 49980 53356 53356 53356 53356 53356 53356 53356		D64006	Synechocystis sp.	238	₽
1216493 1217183 1220068 1220068 1219710 1220630 1221645 1223894 1223894 1223894 1223894 1223894 122628 1226528 1226528 1226528 1226528 1226528 1226528 1226528 17764 77164 79724 88721					
1217183 1220068 1219710 1220630 1221645 1221645 1223894 1225000 1225000 1226528 1226528 1226528 1226972 47569 49980 49980 53356 53477 63753 77164 79724					
1220068 1219710 1220630 1221645 1223894 1223894 1225000 1225000 1226528 1226528 1226528 1226528 1226528 1729972 47569 49980 49980 53356 53477 63753 77164 79724					T
1219710 1220630 1221645 1223894 1223894 1225000 1227810 1226528 1226528 1226528 1226972 47569 49980 53356 53477 63753 77164 79724 88721					
1220630 1221645 1223894 1225800 1225800 1226528 1226972 47569 49980 53356 53477 63753 77164 79724 88721					
1221645 1223894 1225000 1225000 1226528 1229972 47569 49980 53356 54477 63753 77164 79724 88721	376 putative			;	;
1223894 1225000 1225000 1226528 1229972 47569 49980 53356 53477 63753 77164 79724 88721	581 unknown	D26185	Bacillus subtilis	179	54
1225000 1227810 1226528 1229972 47569 49980 53356 53477 63753 77164 79724 88721	988 putative			30,	Ţ
1227810 1226528 1229972 47569 49980 53356 54477 63753 77164 79724 88721	330 high level kasgamycin resistance	D26185	Bacillus subtilis	477	4
1226528 1229972 47569 49980 53356 53477 63753 77164 79724 88721		D90903	Synechocystis sp.	1129	54
1229972 1 47569 4 49980 4 53356 5 53477 5 63753 6 77164 7 79724 8					Ţ
47569 49980 53356 54477 63753 77164 79724 88721	_	U32723	Haemophilus influenzae	999	9 5
53356 53477 63753 77164 79724 88721		AE001308	Chlamydia trachomatis	/10	7/
53356 54477 63753 77164 79724 88721					
54477 63753 77164 79724 88721 94067				211	7
63753 77164 79724 88721 94067		AE001307	Chlamydia trachomatis	311	
77164 79724 88721 94067		AE001306	Chlamydia trachomatis	198	10
79724 88721 94067	7 putative			97,	5
88721 94067		AE001302	Chlamydia trachomatis	428	ò
94067			1		
				300	
		AE001303	Chlamydia trachomatis	3%0	5
ORF1148 147536 147234	34 putative				

Cationic Amino Acid Transporter
Cationic Amino Acid Transporter
Dicarboxylate Translocator
Disulfide bond Oxidoreductase

ORF	Begin	End	Homology	QI	Species	Score	
							,
ORF1182	337348	338289	sodium-dependent transporter	AF017105	Chlamydia psittaci	7111	77
ORF1183	364764	364369	Prolipoprotein Diacylglycerol Transferase	AE001298	Chlamydia trachomatis		4
ORF1184	389623	390135	hypothetical protein	AE001282	Chlamydia trachomatis	2	3
ORF1185	393729	394343	ABC superfamily ATPase	AE001282	Chlamydia trachomatis	4/3	25
ORF1186	407379	407621	putative				
ORF1187	410944	410708	putative				
ORF1188	427632	427988	putative				
ORF1189	428172	428486	putative				;
ORF1190	436761	437246	hypothetical protein	AE001279	Chlamydia trachomatis	199	81
ORF1191	460911	461159	putative				-
ORF1192	477597	477313	hypothetical protein	AE001300	Chlamydia trachomatis	309	79
ORF1193	487303	487001	putative				7
ORF1194	487764	487534	Glycine Cleavage System H Protein	AE001300	Chlamydia trachomatis	221	6
ORF1195	498502	499017	hypothetical protein	AE001275	Chlamydia trachomatis	206	32
ORF1196	499795	500466	putative				
ORF1197	571928	572344	putative				
ORF1198	572367	572131	putative			1,36	(
ORF1199	588184	587915	hypothetical protein	AE001312	Chlamydia trachomatis	200	70
ORF1200	600587	206009	(Metalloenzyme)	AE001316	Chlainydia trachomatis	314	7
ORF1201	609731	608895	putative			1	ļ
ORF1202	614039	614755	hypothetical protein	AE001317	Chlamydia trachomatis	4/5	9
ORF1203	614823	615152	putative			;	1;
ORF1204	638244	638831	ABC Transporter ATPase	AE001315	Chlamydia trachomatis	614	10
ORF1205	638819	639094	(Metal Transport Protein)	AE001315	Chlamydia trachomatis	265	63
ORF1206	639073	639636	(Metal Transport Protein)	AE001315	Chlamydia trachomatis	/80	6
ORF1207	647901	648236	hypothetical protein	AE001317	Chlamydia trachomatis	139	×
ORF1208	678510	679469	phosphohydrolase	AE001320	Chlamydia trachomatis	995	63
ORF1209	688178	688732	hypothetical protein	AE001320	Chlamydia trachomatis	366	43
ORF1210	696045	696563	methyltransferase	AE001321	Chlamydia trachomatis	369	49
ORF1211	708998	708588	Glucose-1-P Adenyltransferase	AE001322	Chlamydia trachomatis	507	83
ORF1212	208608	710089	putative			į	ļ
ORF1213	718240	717737	Glycerol-3-P Phosphatidyltransferase	AE001323	Chlamydia trachomatis	5/3	8 2
ORF1214	737828	737565	S19 Ribosomal Protein	AE001323	Chlamydia trachomatis	439	*

ORF	Begin	End	Homology	ar	Species	Score	~ %
						,	19
ORE1215	779502	780257	hypothetical protein	AE001322	Chlamydia trachomatis	476	8
ORF1216	806310	805864	hypothetical protein	AE001337	Chlamydia trachomatis	512	29
ORF1217	820931	820707	putative			1,3	Ş
ORF1218	837696	839096	Exodeoxyribonuclease V, Gamma	AE001334	Chlamydia trachomatis	7967	49
ORF1219	883307	883549	putative				
ORF1220	892010	891726	putative				
ORF1221	893277	893564	putative			130	1
ORF1222	936998	937225	Gen. Secretion Protein E	AE001327	Chlamydia trachomatis	907	
ORF1223	946865	947419	putative			5,5	1
ORF1224	975187	975411	SWF/SNF family helicase	AE001341	Chlainydia trachomatis	203	2 2
ORF1225	985882	985517	hypothetical protein	AE001342	Chlainydia trachomatis	991	2
ORF1226	987713	987180	hypothetical protein	AE001342	Chlamydia trachomatis	447	59
OPE1227	088215	987733	Flagellar M-Ring Protein	AE001342	Chlamydia trachomatis	304	4
OP F1227	088754	988530	Flagellar M-Ring Protein	AE001342	Chlainydia trachomatis	92	36
Op 51730	002547	992841	hymothetical protein	AE001343	Chlamydia trachomatis	112	39
ODE1220	25,750	993067	hymothetical protein	AE001343	Chlamydia trachomatis	100	32
OBE1231	1004247	1004528	D-Ala/Glv Permease	AE001344	Chlamydia trachomatis	283	8
OPE1232	1015013	1014294	235aa long hypothetical protein	AB009472	Pyrococcus horikoshii	104	54
OP 51733	1056147	1056545	putative				
ORF1234	1077682	1078035	predicted disulfide bond isomerase	AE001351	Chlamydia trachomatis	233	46
OPE1235	1088121	1088381	putative				
OPE1236	1098430	1098852	Predicted Kinase	AE001352	Chlamydia trachomatis	384	29
ORF1237	1098798	1099319	Predicted Kinase	AE001352	Chlamydia trachomatis	322	45
ORF1238	1123198	1123515	Transport Permease	AE001354	Chlamydia trachomatis	313	72
ORF1239	1123606	1124256	Tyrosine Transport	AE001354	Chlamydia trachomatis	577	28
ORF1240	1124453	1124797	Tyrosine Transport	AE001354	Chlamydia trachomatis	323	3
ORF1241	1129253	1129567	putative				1
ORF1242	1164947	1164474	hypothetical protein	AE001357	Chlamydia trachomatis	412	3
ORF1243	1170457	1170053	hypothetical protein	AE001358	Chlamydia trachomatis	283	5
ORF1244	1172342	1171863	ABC transporter permease	AE001358	Chlamydia trachomatis	457	2
ORF1245	1192155	1192835	putative				
ORF1246	1192759	1192992	putative				
ORF1247	1193861	1194142	putative				7

ORF	Begin	End	Homology	<b>GII</b>	Species	Score	%1
000000	1104036	1102770	M. Amino Acid Dehydrogenase)	AE001311	Chlamydia trachomatis	269	79
ORF1249	1209748	1209053	conserved hypothetical protein	AE000958	Archaeoglobus fulgidus	121	38
ORF1250	1215111	1215419	putative				
ORF1251	1216302	1216538	putative				18
ORF1252	1228072	1227818	hypothetical protein	AE001306	Chlamydia trachomatis	134	39
ORF1253	1228304	1228080	xseB	AL021897	Mycobacterium tuberculosis	68	33
ORF1254	26599	26222	putative				
ORF1255	27609	27367	putative				
ORF1256	67206	29699	putative				
ORF1257	70612	70352	putative				
ORF1258	132703	132945	putative				
ORF1259	178073	178393	putative		1		
ORF1260	208576	208349	putative				
ORF1261	209156	208929	putative				
ORF1262	209263	209024	putative				
ORF1263	210304	210639	putative				
ORF1264	299009	299452	putative				
ORF1265	352106	351717	putative				!
ORF1266	420182	419949	Flagellar Secretion Protein	AE001280	Chlamydia trachomatis	115	43
ORF1267	553602	553381	putative		1		
ORF1268	556538	556807	putative				
ORF1269	594348	593797	putative				
ORF1270	595169	594876	putative				
ORF1271	662148	662381	putative				
ORF1272	706528	706893	putative				
ORF1273	803315	803650	putative				
ORF1274	849551	849306	putative		-		
ORF1275	913676	913275	putative				
ORF1276	927087	926836	putative				
ORF1277	930587	930360	putative			,	ç
ORF1278	986531	986764	ORF 12	M72718	Bacillus subtilis	901	48
ORF1279	996229	996486	putative				
ORF1280	1000373	1000002	putative				

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ORF	Begin	End	Homology	<b>a</b>	Species	2000	0/1
OPE1281	1010291	1010037	putative				
ORF1282	1011128	1010793	106aa long hypothetical protein	AB009472	Pyrococcus horikoshii	159	20
ORF1283	1012924	1012694	putative				
ORF1284	1028659	1028913	putative				
ORF1285	1086481	1086762	putative			3	1
ORF1286	1118658	1118879	Phosphoglucomutase	AE001354	Chlamydia trachomatis	291	<b>%</b>
ORF1287	1170098	1169835	hypothetical protein	AE001358	Chlamydia trachomatis	187	53
ORF1288	1180828	1181184	putative				
ORF1289	1182658	1183035	putative				
ORF1290	1195076	1194795	putative				
ORF1291	1195890	1196183	putative				]

Table 2

ORF Nos	begin	end	potential start
2	42	794	42
3	1258	1614	1261
4	1807	2418	1807
5	3393	2491	3393
6	3639	4067	3639
7	5649	4270	5649
8	7463	6012	7463
9	8051	8962	8051
10	9129	9959	9138
. 11	10687	10361	_10639
12	10927	11232	10927
13	11246	12727	11246
14	12691	14190	12691
15	14484	17249	14484
16	16039	15770	16036
17	17845	20853	17845
18	21137	22042	21137
19	22046	23476	22046
20	23681	26110	23681
21	26109	25861	26109
22	26241	26978	26241
23	26960	27754	26960
24	27747	28577	27747
25	28887	29492	28950
26	29432	30028	29432
27	30024	31472	30024
28	31758	32288	31758
29	32201	33991	32201
30	33852	34541	33852
3:	34783	36063	34783
32	36009	37529	36009
3:	3 37881	39362	37881
34	39418	39161	39418

OPENIO	- Lagin I	end	potential start
ORF Nos	begin		
35	39366	40715	39366
36	43076	41094	43076
37	43800	43066	43800
38	44828	43785	44768
39	45340	44753	45340
40	45752	45372	45752
41	46996	45701	46996
42	47961	47569	47961
43	48960	48040	48960
44	51452	50133	51452
45	52606	51335	52606
46	53684	53319	53684
47	54195	53746	54195
48	55278	56453	55278
49	56493	57266	56493
50	57297	58526	57297
51	59851	58565	59851
52	61495	59924	61495
53	61324	62151	61324
54	62132	62470	62132
55	62474	63733	62474
56	63881	64186	63881
57	64611	64318	64611
58	65485	64673	65485
59	65999	65301	65999
60	66244	67281	66244
61	67265	67699	67265
62	67703	68539	67760
63	68805	70736	68805
64	69172	68831	69172
65			70642
60			71325
6			7 72060
61	1		<u>.l.,</u>
			<u> </u>

ORF Nos	begin	end	potential start
			100000000000000000000000000000000000000
69	78351	77680	78351
70	79356	78355	79356
71	79983	79693	79983
72	80441	79938	80441
73	80475	80969	80475
74	81296	83080	81332
75	83291	83932	83291
76	84005	84769	84005
77	84975	85244	84975
78	85123	85425	85123
79	85397	85903	85397
80	85909	86583	85909
81	86626	88065	86626
82	89257	91026	89257
83	91291	93030	91291
84	93295	94086	93295
85	95285	94707	95279
86	95667	96557	95667
87	96317	97456	96317
88	98435	97968	98435
89	99460	98426	99460
90	100144	101325	100144
91	101457	101720	101457
92	101704	102273	101704
93	102356	102805	102356
94	102835	103530	102835
9:	103549	104058	103549
90	104096	104491	104096
9	7 104601	108386	104601
9:	10840	112054	108401
9	9 112033	112590	112033
10	0 112672	113682	112672
10			113726
10		<u> </u>	L
	1	<u> </u>	L

ORF Nos	begin	end	potential start
103	115267	115755	115267
		116543	115911
104	115911		
105	116736	118055	116778
106	117968	118522	117968
107	118530	119843	118530
108	119816	120457	119816
109	120451	122430	120451
110	122504	122950	122504
111	123528	126347	123528
112	126332	129166	126332
113	134690	129213	134690
114	134925	136382	134931
115	137870	136482	137867
116	137899	138240	137899
117	138239	137928	138239
118	139558	138257	139558
119	140352	139516	140352
120	140498	141841	140498
121	141855	142658	141855
122	144258	143050	144258
123	145258	144494	145258
124	145454	146749	145454
125	147318	146767	147318
126	148261	147677	148261
127	149029	152157	149029
128	154108	152201	154108
129	155135	154308	155135
130	155141	155467	155141
131		156779	155703
132	156748	157635	156748
133			
134			<u> </u>
135			
130			

ORF Nos	begin	end	potential start
137	160898	161539	160898
138	161527	162153	161527
139	162144	162443	162144
140	162437	164098	162437
141	165451	164228	165451
142	166349	165411	166349
143	166949	168442	166949
144	169416	171029	169416
145	170857	171459	170857
146	172652	173428	172652
147	174626	173439	174626
148	174816	175613	174816
149	175598	175954	175598
150	175958	176935	175958
151	177708	176938	177708
152	177128	177376	177128
153	179472	177841	179472
154	179822	179517	179822
155	181793	179943	181793
156	182628	181876	182628
157	184420	183074	184420
158	184988	184467	184988
. 159	185483	185112	185483
160	185902	185483	185902
161	186174	185839	186174
162	187720	186587	187720
163	188318	190933	188318
164	191090	191635	191090
165	191547	192743	191547
160	192969	193469	192969
16	7 194044	193610	194044
16	194196	195809	194196
169	9 196088	19807:	196088
170	0 198132	199454	198132

0253			potential start
ORF Nos	begin:	end	
171	199351	202818	199351
172	204552	202999	204552
173	205648	204692	205639
174	205807	207327	205807
175	207182	207775	207182
176	207779	208267	207779
177	208267	209577	208267
178	211807	211271	211807
179	212188	211844	212188
180	214079	212448	214079
181	214907	214083	214907
182	216154	215429	216154
183	216115	216678	216115
184	216728	217282	216728
185	217267	217866	217267
186	218593	218261	218590
187	219821	218994	219821
188	221382	220309	221382
189	222719	221433	222719
190	223521	222724	223521
191	224499	225008	224499
192	225140	225559	225140
193	225555	226802	225555
194	227800	226892	227743
195	228335	228072	228335
196	229251	228643	229251
197	230983	229622	230983
198	231483	230983	231483
199	232063	231509	232063
200	232739	232053	232739
20	233166	234356	233166
200	2 233518	233165	233518
20	3 234536	235186	234536
20	4 235379	236689	235379
		1	L

ORF Nos	begin	end	potential start
205	236680	237618	236689
206	237521	238345	237521
207	238281	238973	238281
208	238871	240115	238871
209	240191	241564	240191
210	242281	241604	242281
211	242933	242274	242933
212	243416	242976	243416
213	243500	244531	243500
214	244480	246021	244480
215	246330	247811	246330
216	247831	249174	247870
217	249437	251038	249455
218	251325	252212	251325
219	253156	254007	253156
220	253974	254852	253974
221	255258	256094	255258
222	256640	257455	256640
223	257502	258239	257502
224	257869	257501	257869
225	259248	260897	259248
226	262753	261788	262753
227	263059	262757	263059
228	264375	263182	264375
229	265985	264747	265985
230	266637	266059	266637
23	267338	266538	267338
232	2 267922	267473	267922
23:	269647	27077	269647
234	272777	273145	272777
23:	273253	273636	273253
23	6 273705	27397	7 273705
23	7 276016	27571	7 276016
23	8 276439	27602	276418

239			potential start
	276792	277253	276792
240	277318	277599	277318
241	278578	277877	278578
242	279258	278554	279258
243	280435	279533	280435
244	281547	280849	281547
245	281696	282325	281717
246	282459	284069	282459
247	284056	284517	284056
248	284606	285775	284606
249	285592	285987	285592
250	286179	286976	286179
251	287583	287002	287583
252	287951	287451	287951
253	288499	288816	288499
254	289674	288505	289674
255	288839	289213	288839
256	289970	290254	289970
257	291931	292803	291931
258	293258	292755	293258
259	293718	293272	293718
260	294630	293953	294630
261	296153	294636	296153
262	294817	295068	294817
263	296354	297862	296354
264	298415	297879	298415
265	298777	298253	298777
266	299572	298781	299572
267	300487	299633	300487
268	301586	300702	301568
269	302440	301571	302440
270	302838	302437	302838
271	303335	302745	303335
272	304394	303852	304394

273         304606         305223         304606           274         305394         306236         305394           275         306501         307439         306501           276         308033         307458         308033           277         308924         308037         308924           278         309485         310180         309485           279         310426         311214         310426           280         311597         311253         311504           281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291 <th>ORF Nos</th> <th>begin</th> <th>end</th> <th>potential start</th>	ORF Nos	begin	end	potential start
275         306501         307439         306501           276         308033         307458         308033           277         308924         308037         308924           278         309485         310180         309485           279         310426         311214         310426           280         311597         311253         311504           281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323856         324878           294 <td></td> <td></td> <td>305223</td> <td>304606</td>			305223	304606
276         308033         307458         308033           277         308924         308037         308924           278         309485         310180         309485           279         310426         311214         310426           280         311597         311253         311504           281         312772         313780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323856         324878           293         324878         323856         324878           294 <td>274</td> <td>305394</td> <td>306236</td> <td>305394</td>	274	305394	306236	305394
277         308924         308037         308924           278         309485         310180         309485           279         310426         311214         310426           280         311597         311253         311504           281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295 <td>275</td> <td>306501</td> <td>307439</td> <td>306501</td>	275	306501	307439	306501
278         309485         310180         309485           279         310426         311214         310426           280         311597         311253         311504           281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296 <td>276</td> <td>308033</td> <td>307458</td> <td>308033</td>	276	308033	307458	308033
279         310426         311214         310426           280         311597         311253         311504           281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297 <td>277</td> <td>308924</td> <td>308037</td> <td>308924</td>	277	308924	308037	308924
280         311597         311253         311504           281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298 <td>278</td> <td>309485</td> <td>310180</td> <td>309485</td>	278	309485	310180	309485
281         312772         311780         312772           282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299 <td>279</td> <td>310426</td> <td>311214</td> <td>310426</td>	279	310426	311214	310426
282         313425         312772         313425           283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330556         332455           301 <td>280</td> <td>311597</td> <td>311253</td> <td>311504</td>	280	311597	311253	311504
283         313646         313377         313646           284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301 <td>281</td> <td>312772</td> <td>311780</td> <td>312772</td>	281	312772	311780	312772
284         313937         314665         313937           285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301         336091         334877         336091           302 <td>282</td> <td>313425</td> <td>312772</td> <td>313425</td>	282	313425	312772	313425
285         315576         314755         315576           286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323856         324878           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301         336091         334877         336091           302         336103         337302         336103           303 <td>283</td> <td>313646</td> <td>313377</td> <td>313646</td>	283	313646	313377	313646
286         316157         315531         316157           287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301         336091         334877         336091           302         336103         337302         336103           303         338129         338830         338129           304 <td>284</td> <td>313937</td> <td>314665</td> <td>313937</td>	284	313937	314665	313937
287         318657         316156         318657           288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301         336091         334877         336091           302         336103         337302         336103           303         338129         338830         338129           304         338965         339501         338965           305 <td>285</td> <td>315576</td> <td>314755</td> <td>315576</td>	285	315576	314755	315576
288         321042         318676         321042           289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301         336091         334877         336091           302         336103         337302         336103           303         338129         338830         338129           304         338965         339501         338965           305         339508         340143         339508	286	316157	315531	316157
289         321445         321098         321445           290         322309         321710         322309           291         323190         322366         323181           292         323843         323181         323843           293         324878         323856         324878           294         325340         326410         325340           295         326433         327836         326433           296         328465         327839         328465           297         329360         328857         329360           298         330907         329357         330907           299         332455         330956         332455           300         334536         332395         334536           301         336091         334877         336091           302         336103         337302         336103           303         338129         338830         338129           304         338965         339501         338965           305         339508         340143         339508	287	318657	316156	318657
290       322309       321710       322309         291       323190       322366       323181         292       323843       323181       323843         293       324878       323856       324878         294       325340       326410       325340         295       326433       327836       326433         296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	288	321042	318676	321042
291       323190       322366       323181         292       323843       323181       323843         293       324878       323856       324878         294       325340       326410       325340         295       326433       327836       326433         296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	289	321445	321098	321445
292       323843       323181       323843         293       324878       323856       324878         294       325340       326410       325340         295       326433       327836       326433         296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	290	322309	321710	322309
293       324878       323856       324878         294       325340       326410       325340         295       326433       327836       326433         296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	291	323190	322366	323181
294       325340       326410       325340         295       326433       327836       326433         296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	292	323843	323181	323843
295       326433       327836       326433         296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	293	324878	323856	324878
296       328465       327839       328465         297       329360       328857       329360         298       330907       329357       330907         299       332455       330956       332455         300       334536       332395       334536         301       336091       334877       336091         302       336103       337302       336103         303       338129       338830       338129         304       338965       339501       338965         305       339508       340143       339508	294	325340	326410	325340
297     329360     328857     329360       298     330907     329357     330907       299     332455     330956     332455       300     334536     332395     334536       301     336091     334877     336091       302     336103     337302     336103       303     338129     338830     338129       304     338965     339501     338965       305     339508     340143     339508	295	326433	327836	326433
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299     332455     330956     332455       300     334536     332395     334536       301     336091     334877     336091       302     336103     337302     336103       303     338129     338830     338129       304     338965     339501     338965       305     339508     340143     339508	297	329360	328857	329360
300     334536     332395     334536       301     336091     334877     336091       302     336103     337302     336103       303     338129     338830     338129       304     338965     339501     338965       305     339508     340143     339508	298	330907	329357	330907
301     336091     334877     336091       302     336103     337302     336103       303     338129     338830     338129       304     338965     339501     338965       305     339508     340143     339508	299	332455	330956	332455
302     336103     337302     336103       303     338129     338830     338129       304     338965     339501     338965       305     339508     340143     339508	300	334536	332395	334536
303     338129     338830     338129       304     338965     339501     338965       305     339508     340143     339508	301	336091	334877	336091
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305 339508 340143 339508	303	338129	338830	338129
	304	338965	339501	338965
306 340247 342967 340247	305	339508	340143	339508
	306	340247	34296	340247

ORF Nos	begin	end	potential start
307	343385	343810	343385
308	344171	343935	344171
309	345082	344330	345073
310	346005	345082	346005
311	346784	346437	346784
312	347029	346715	347029
313	347034	347723	347034
314	348075	350459	348075
315	350598	351071	350598
316	351075	352175	351096
317	353291	352230	353267
318	353442	354467	353442
319	354451	354933	354451
320	355000	355449	355000
321	355448	356743	355448
322	355953	355642	355953
323	359310	356827	359310
324	359120	359377	359120
325	359525	359908	359525
326	361290	359947	361290
327	363785	361362	363746
328	364496	363888	364496
329	364832	365290	364832
330	365304	365669	365304
331	366599	365667	366599
332	367291	369030	367291
333	369134	369808	369134
334	369917	370438	369917
335	370365	372647	370365
330	372557	373066	372557
33	7 373020	373442	373020
338	8 373467	374195	373467
339	9 374176	375099	374176
340	0 375676	375083	375676

			2007 3207 224 0 <b>1</b> 024 322000
ORF Nos	begin	end	potential start
341	376173	375634	376173
342	376564	377643	376564
343	377956	379773	377956
344	379781	380425	379805
345	380281	381000	380281
346	381008	381460	381008
347	381460	383037	381460
348	383257	383523	383257
349	383553	385304	383553
350	385397	386458	385400
351	387242	386514	387242
352	388764	387013	388764
353	390120	390932	390120
354	390919	391818	390961
355	392379	391885	392379
356	392582	392986	392582
357	392776	393684	392776
358	394151	394804	394151
359	394928	395308	394928
360		395990	395259
361		395953	397815
362		397831	398850
363		399099	400085
364		400073	401236
365	401474	401136	401474
366			
367			
368			
369			
370			
37			
37:			
37:		<u> </u>	
37.			
31.	712702	, 41702	712.02

ORF Nos	begin	end	potential start
375	415402	414407	415402
376	415848	415237	415848
377	417131	415866	417131
378	417258	417566	417258
379	418326	417454	418326
380	420057	418426	420057
381	420448	420720	420448
382	420980	421552	420980
383	421556	422029	421556
384	422461	422925	422461
385	423562	424320	423562
386	424250	424591	424250
387	424830	426047	424830
388	426240	427397	426240
389	428841	430703	428841
390	430694	431446	430694
391	431597	432100	431597
392	432165	432779	432165
393	433272	432832	433272
394	433925	433227	433922
395	436678	433934	
396	437176	438357	437176
397	440317	438518	440317
398	440001	440345	440001
399	441233	440517	441233
400	440719	441012	440719
401	442192	441230	442192
402	2 442888	442343	442888
403	3 442371	442961	442371
404	4 443578	443003	443578
40:	5 444500	443526	444500
400	6 444842	444528	3 444842
40	7 445009	444743	3 445009
40	8 445718	445182	445718
			<del></del>

ORF Nos	begin	end	potential start
409	445807	447804	445807
410	448738	447803	448738
411	449628	448618	449628
412	450298	. 450867	450298
413	450713	451207	450713
414	451211	452452	451211
415	452448	453659	452448
416	454843	453725	454843
417	455608	454865	455608
418	456243	457007	456243
419	457016	457708	457016
420	458368	457979	458368
421	459496	458372	459496
422	459493	460194	459493
423	461446	460355	461446
424	462298	461450	462298
425	462444	463349	462444
426	464241	463342	464241
427	464574	465065	464574
428	465129	465611	465129
429	465571	466317	465571
430	466317	467093	466317
431	466999	467502	466999
432	469691	467715	469691
433	470691	469660	470691
434	472010	470709	472010
435	471545	471799	471545
436	472359	472045	472359
437	473523	472732	473523
438	474889	473441	474889
439	477323	475365	477323
440	478496	477597	478496
441	478722	479273	478722
442	479277	479705	479277

ORF Nos	begin	end	potential start
443	480050	481450	480050
444	481469	482053	481469
445	482600	482025	482600
446	482654	484204	482654
447	484211	485170	484211
448	485170	485838	485170
449	485813	486580	485813
450	486976	486638	486976
451	489071	487764	489071
452	489341	489090	489341
453	489958	489152	489958
454	490549	489962	490549
455	491163	490522	491163
456	491396	491112	491396
457	492121	491390	492121
458	492304	494838	492304
459	495943	494822	495943
460	496011	496565	496170
461	496569	497228	496569
462	497358	497834	497358
463	497770	498327	497770
464	499209	499589	499209
465	499520	499792	499520
466	500774	504169	500774
467	504139	504600	504139
468	504865	506877	504865
469	506790	507671	506790
470	507718	510507	507718
471	508325	507912	508325
472	510660	513440	510660
473	514965	513787	514920
474	517347	515419	517347
475	517058	517363	517058
470	517798	517277	517798

ORF Nos	begin	end.	potential start
477	518200	517847	518200
478	518300	521146	518363
479	521392	522948	521407
480	523244	524809	523322
481	524379	524125	524379
482	524649	526238	524649
483	526265	527104	526268
484	526947	526702	526947
485	526975	528450	526975
486	528408	529199	528408
487	530612	529542	530612
488	531656	530616	531656
489	533974	532067	533974
490	536432	534324	536432
491	537150	536707	537150
492	537928	537080	537928
493	538438	537932	538438
494	538737	538333	538737
495	539594	539127	539594
496	541215	539590	541215
497	542571	541282	542571
498	543014	542457	543014
499	543369	542962	543369
500	543809	546628	543815
501	546619	549525	546619
502	547293	546994	547293
503	549699	550523	549699
504	550490	551551	550490
503	551448	552623	551448
500	5 552652	555117	552652
50	555029	555493	555029
508	558006	55567	558006
509	559694	558162	559694
510	558208	55857	558208

ORF Nos	begin	end	potential start
511	561692	559899	561692
512	561412	561708	561412
513	563942	561777	563942
514	564969	563950	564969
515	566204	564936	566198
516	567717	566302	567717
517	568526	567708	568526
518	569467	568742	569467
519	571065	569431	571065
520	571828	571118	571783
521	572202	573308	572202
522	573146	575056	573146
523	575023	575916	575023
524	577891	576497	577891
525	578914	578204	578914
526	579924	578857	579924
527	580187	579858	580187
528	580017	580406	580017
529	581086	580187	581086
530	581367	581828	581367
531	581678	582367	581678
532	582361	583428	582361
533	584690	583431	584690
534	585237	584950	585237
535	585626	586888	585626
536	586846	58790	586888
537	589049	588180	589049
538	590500	58930	590455
539	59075	59245	590755
540	592520	59290	3 592526
54	59283	59374	7 592836
542	59374	7 59429	8 593747
543	3 59433	1 59594	7 594331
54	4 59590	5 59630	9 595905

ORF Nos	begin	end	potential start
545	596514	597215	596514
546	597184	597957	597184
547	597755	598612	597755
548	598602	599204	598602
549	599373	599939	599373
550	600903	602072	600903
551	602240	602587	602240
552	602637	603272	602637
553	603142	604512	603142
554	604627	605853	604627
555	605790	606620	605790
556	606571	607281	606571
557	609004	607355	609004
558	610906	609932	610906
559	611786	611004	611786
560	612333	611746	612333
561	613897	612341	613897
562	615179	616279	615179
563	616610	617383	616610
564	618796	617810	618796
565	620004	618826	620004
566	619649	619918	619649
567	621265	620021	621265
568	622359	621265	622359
569	623420	622560	623420
570	624297	623335	624297
571	624773	624174	624773
572	625029	625484	625029
573	625488	625883	625488
574	625892	626395	625892
575	626444	627790	626444
576	627912	628607	627930
577	628774	629697	628774
578	629660	631639	629660

ORF Nos	begin	end	potential start
579	631725	633551	631725
580	633520	636957	633520
581	637232	638098	637232
582	640648	639593	640648
583	640979	640728	640979
584	641327	641007	641327
585	641687	642283	641687
586	643023	642286	643023
587	643330	643076	643330
588	643704	643351	643704
589	645628	643676	645628
590	645783	645538	645756
591	646269	645793	646269
592	646751	646314	646751
593	647848	647045	647848
594	648393	650336	648393
595	651016	650420	651007
596	652956	651289	652956
597	653395	653126	653395
598	655740	654193	655740
599	656508	655966	656508
600	658140	657022	658140
601	660216	658525	660216
602	663238	660248	663238
603	664461	663157	664452
604	665735	664635	665735
605	666212	666994	666212
600	666998	66792	666998
601	667909	668568	667909
608	668502	669203	668502
609	669154	670893	669175
610	67222	67085	672226
61	67113	67142	671137
61:	67245	67300	1 672453

ORF Nos	begin	end	potential start
613	673072	674721	673072
614	674549	674262	674549
615	675518	674796	675518
616	676083	675499	676083
617	676630	676067	676630
618	677016	676600	677016
619	677647	677015	677647
620	677990	678259	677990
621	679444	680097	679444
622	680097	680897	680097
623	681637	680849	681637
624	681409	682281	681409
625	682453	682821	682453
626	682763	683902	682763
627	684616	683969	
628	685169	684534	685169
629	685986	685117	685986
630	686278	687288	686278
631	687483	688151	687483
632	688740	689501	688740
633	690242	689622	
634	690470	691126	
635	692600	691497	
630	692674	695064	<u> </u>
63	695049	696032	
638	697964	696585	697964
639	699803	698274	
64	701926	69978	
64	703190	70256	
64			
64	70424	1	
64	4 706070	70530	706070
64	5 70684	70625	
64	6 70759	70681	707596

ORF Nos	begin	end	potential start
647	708666	707677	708666
648	709793	709119	709793
649	711523	710132	711523
650	712236	711523	712236
651	714734	712125	714734
652	715759	714761	715759
653	717538	715886	717538
654	719113	720243	719113
655	720590	722422	720590
656	722406	723056	722406
657	723551	723120	723551
658	724246	723626	724246
659	724754	724251	724754
660	725868	724900	725868
661	727115	726270	727115
662	728126	727119	728126
663	728594	728208	728594
664	729614	728604	729614
665	729778	729533	729778
666	730149	729751	730149
667	730539	730174	730539
668	731983	730598	731983
669	732427	731996	732427
670	732917	732423	732917
671	733598	733320	733598
. 672	733869	733492	733869
673	734298	733900	734298
674	734858	734319	734858
675	735195	734863	735195
676	735578	735342	735578
677	735861	735604	735861
678	736492	736079	736492
679	737192	736524	737192
680	737555	737211	737555

ORF Nos	begin	end	potential start
681	738688	737837	738688
682	739048	738713	739048
683	739736	739065	739736
684	740477	739773	740477
685	740659	740958	740659
686	741722	740721	741722
687	742789	741827	742789
688	743618	742782	743618
689	744092	743634	744092
690	744604	744107	744604
691	744953	744498	744953
692	746608	744986	746608
693	747085	746621	747085
694	747974	747219	747974
695	748594	748169	748594
696	749145	748573	749145
697	749652	749957	749652
698	750446	749979	750446
699	751219	750446	751219
700	753042	751291	753042
701	754309	753020	754309
702	755120	756175	755120
703	756120	756485	756120
704	756499	760227	756499
705	761217	760297	761178
706	761297	761809	761330
707	761782	762282	761782
708	762260	762895	762299
709	762867	763316	762867
710	763780	763325	763780
711	763861	765168	763861
712	766809	765691	7 766809
713	768051	766888	768051
714	768566	76832	768566

ORF Nos	begin	end	potential start
715	769342	768551	769342
716	770532	769378	770532
717	771451	770804	771451
718	773058	771847	773058
719	773094	773456	773094
720	774376	773093	774376
721	775123	774380	775123
722	775398	774916	775398
723	775046	776077	775046
724	776070	777041	776070
725	777964	777536	777964
726	778176	777904	778176
727	778621	779334	778684
728	781173	780307	781173
729	781526	781116	781526
730	782784	781555	782784
731	783572	782805	783572
732	785032	783581	785032
733	786412	785360	786412
734	788429	786450	788429
735	788944	788528	788944
736	789758	788901	789758
737	790332	791504	790338
738	791846	792721	791846
739	792724	793569	792724
740	793580	794323	793580
741	794304	794843	794304
742	795217	795732	795217
743	795722	796795	795722
744	798735	797053	798735
745	799823	798681	799823
746	799297	799578	799297
747	801313	799808	801313
748	802453	801332	802453

ORF Nos	begin	end	potential start
749	803299	802457	803299
750	803811	803290	803811
751	805151	803826	805151
752	805860	805156	805860
753	806604	806332	806604
754	806913	806608	806913
755	808222	806903	808222
756	808751	808146	808751
757	809437	808673	809437
758	809939	809454	809939
759	811235	810213	811235
760	811779	813056	811779
761	812890	812516	812890
762	812954	813583	812954
763	813587	815023	813587
764	815420	815746	815420
765	816036	817010	816036
766	817111	817356	817111
767	817791	818609	817797
768	818609	819094	818609
769	819104	819823	819104
770	820722	819826	820722
771	822313	821000	822313
772	823503	822238	823503
773	823678	825612	
774	825461	826312	825461
775	827280	82664	
776	828604	82717	828604
777	830026	82871	
778	831047	83008	
779	831725	83105	831725
780	832220	83309	8 832220
78	833851	83339	_l
783	834068	83503	9 834068

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ORF Nos	begin	end	potential start
783	835792	835127	835792
784	837624	836116	837624
785	838951	840882	838951
786	840869	842185	840869
787	841989	843455	841989
788	843242	844021	843242
789	845018	843987	844997
790	846174	844990	846174
791	848509	846311	848509
792	848568	849014	848568
793	849082	850488	849088
794	851512	850574	851512
795	852064	852447	852064
796	852398	853690	852398
797	855118	854243	855118
798	855751	855128	855751
799	856551	855829	856551
800	856730	858556	856730
801	858717	859601	858717
802	859591	860205	859591
803	861132	860284	861132
804	861426	861163	861426
805	861701	862921	861701
806	863026	864798	863026
807	864831	865256	864831
808	865226	866581	865226
809	866562	867119	866562
810	867025	867816	867025
811	867820	868497	867820
812	869743	868661	869743
813	870633	870094	870633
814	871929	870646	871929
815	872538	872086	872538
816	873908	872517	873908
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ORF Nos	begin	end	potential start
817	874281	874670	874281
818	874582	875286	874582
819	877857	875377	877857
820	878446	879255	878446
821	880635	879268	880635
822	882524	880593	882524
823	882612	883319	882612
824	884155	883538	884155
825	884340	885611	884343
826	885722	887302	885722
827	887587	888153	887587
828	888627	888220	888627
829	889330	888716	889330
830	889898	889323	
831	891190	889898	891190
832	891828	891247	
833	892421	892017	
834	893116		
835	892521		
836	893392		
837	895745		<u> </u>
838			
839		<u> </u>	<u> </u>
840			
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85	0 91230	3 91282	1 912303

ORF Nos	begin	end	potential start
	912937	913983	912937
851			
852	915128	914067	915128
853	916658	915303	916658
854	915627	915376	915627
855	917707	916853	917707
856	918837	917722	918837
857	919868	918837	919868
858	920434	919880	920434
859	921187	920438	921187
860	921959	921195	921959
861	923773	921995	923773
862	922146	922415	922146
863	923943	923674	923943
864	924077	925006	924077
865	925436	925083	925436
866	926524	925349	926524
867	927920	926433	927920
868	928319	927951	928319
869	928963	928334	928963
870	929248	930987	929248
871	930995	932059	930995
872	932121	933515	
873		932513	
874			
875		937082	
876		938410	
877		938805	
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880			
881			
882			
883			
884	942784	942500	942763

ORF Nos	begin	end	potential start
885	943149	942799	943149
886	943799	943029	943799
887	944055	943732	944055
888	944413	943994	944404
889	945395	944556	945395
890	945853	945389	945853
891	946392	945751	946392
892	947410	948081	947431
893	949871	948915	949871
894	951058	949868	951058
895	951249	950959	951249
896	951664	952134	951664
897	952674	952165	952674
898	953491	952589	953491
899	955324	953495	955324
900	955823	955281	955823
901	957082	955847	957082
902	957902	957270	957902
903	959231	957906	959231
904	959376	960284	959376
905	960266	961669	960347
906	961856	964765	961856
907	966855	965395	966855
908	968204	966975	968204
909	968791	968237	968791
910	969498	968731	969498
911	969858	969511	969858
912	970118	969762	970118
913	970593	970300	970593
914	971261	970542	971261
91:	971680	971123	971680
910	971876	975100	971876
91	975419	976516	975419
918	976584	978320	976584

ORF Nos	begin	end	potential start
919	977680	977231	977680
920	978399	980738	978399
921	980756	981928	980756
922	982974	981931	982962
923	984120	983119	984120
924	985502	984120	985502
925	987180	985882	987180
926	987172	987444	987172
927	989846	989049	989846
928	991048	989846	991048
929	991638	990955	991638
930	991794	992498	991794
931	993619	993041	993619
932	993530	994792	993548
933	995970	994795	995970
934	996857	995739	996857
935	997603	996782	997603
936	998969	997572	998969
937	998896	1000023	998896
938	1000087	1001340	1000087
939	1001357	1001818	1001357
940	1003288	1001873	1003288
941	1003487	1004146	1003496
942	1004485	1005639	1004689
943	1005643	1005972	1005643
944	1006784	1006116	1006784
945	1007563	1006769	1007563
946	1009226	1007568	1009226
947	1009989	1009336	1009989
948	1015852	1016337	1015852
949	1016561	1016181	1016561
950	1016297	1017532	1016297
951	1016802	1016452	1016802
952	1018993	1017701	1018993
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ORF Nos	1.00	end	potential start
	begin		
953	1019454	1019137	1019454
954	1020764	1019562	1020764
955	1021405	1021037	1021405
956	1021821	1024286	1021821
957	1024697	1024248	1024697
958	1025569	1024508	1025551
959	1026969	1025590	1026969
960	1027789	1026947	1027789
961	1031199	1027945	1031199
962	1031717	1031172	1031717
963	1033057	1031612	1033057
964	1033425	1033039	1033425
965	1033784	1033200	1033784
966	1033963	1036038	1033963
967	1036945	1036010	1036945
968	1037110	1037679	1037110
969	1037696	1037944	1037696
970	1038916	1037975	1038916
971	1040582	1039026	1040582
972	1040997	1042337	1040997
973	1042357	1043403	1042357
974	1043367	1044623	1043367
975	1044607	1045362	1044607
976	1045384	1046538	1045384
977	1046447	1047517	1046447
978	1047521	1049956	1047521
979	1050611	1050036	1050611
980	1050925	1050566	1050925
981	1051728	1051090	1051728
982	1051743	1052063	1051743
983	1052101	1053126	1052101
984	1054201	1053107	1054201
985	1054242	1055555	1054242
986	1055483	1055908	1055483
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ORF Nos	begin	end	potential start
987	1056609	1056965	1056609
988	1056961	1058232	1056985
989	1058238	1058687	1058238
990	1059371	1058727	1059371
991	1059526	1060578	1059526
992	1061553	1060579	1061553
993	1061674	1062411	1061674
994	1062377	1064077	1062377
995	1064116	1065243	1064116
996	1067451	1065178	1067451
997	1068065	1067376	1068065
998	1068209	1068706	1068230
999	1069958	1068819	1069958
1000	1071163	1070033	1071163
1001	1072438	1071332	1072438
1002	1072997	1073476	1072997
1003	1074239	1075864	1074239
1004	1076790	1075867	1076790
1005	1077268	1076573	1077268
1006	1077999	1078724	1077999
1007	1079088	1078672	1079088
1008	1079642	1079944	1079642
1009	1080501	1079995	1080468
1010	1080775	1081341	1080775
1011	1083158	1081350	1083158
1012	1084677	1083235	1084677
1013	1085648	1084632	1085648
1014	1086117	1086737	1086117
1015	1086692	1087897	1086692
1016	1088646	1089005	1088646
1017	1089146	108980	1089146
1018	1092931	1089890	1092931
1019	1093179	1092889	1093179
1020	1093584	1094204	1093584

ORF Nos	begin	end	potential start
1021	1095619	1094192	1095619
1022	1096074	1096628	1096074
1023	1096633	1097082	1096633
1024	1097266	1097601	1097266
1025	1097622	1097867	1097622
1026	1097886	1098392	1097886
1027	1099521	1099279	1099521
1028	1099689	1101053	1099704
1029	1102192	1101107	1102192
1030	1104950	1102116	1104950
1031	1106508	1104946	1106508
1032	1106722	1107249	1106722
1033	1107463	1108101	1107463
1034	1108041	1108421	1108041
1035	1108520	1113370	1108520
1036	1114958	1113447	1114958
1037	1116915	1115071	1116915
1038	1118183	1116894	1118183
1039	1118846	1120030	1118846
1040	1120040	1120522	1120040
1041	1120510	1121430	1120510
1042	1121321	1121866	1121321
1043	1122123	1122899	1122123
1044	1124842	1125564	1124842
1045	1126526	1125579	1126526
1046	1126519	1127676	1126519
1047	1127672	1128571	1127672
1048	1130230	1131336	1130230
1049	1131480	1132553	1131480
1050	1132830	1133843	1132830
105	1134121	1134855	1134121
1052	1134642	1135592	1134642
105	1135964	1135653	1135964
1054	1137132	1135954	1137132

ORF Nos	begin	end	potential start
1055	1137169	1140102	1137169
1056	1141365	1140112	1141344
1057	1142150	1141356	1142150
1058	1142520	1145660	1142520
1059	1145627	1146721	1145627
1060	1146862	1147545	1146862
1061	1147666	1148190	1147666
1062	1148514	1148224	1148514
1063	1149136	1148348	1149136
1064	1149702	1149166	1149702
1065	1150031	1150591	1150031
1066	1150785	1151147	1150785
1067	1151165	1152181	1151165
1068	1152522	1154591	1152522
1069	1155666	1154566	1155666
1070	1156743	1155670	1156740
1071	1156859	1157815	1156859
1072	1157982	1160735	1157982
1073	1162620	1160917	1162620
1074	1162970	1162590	1162970
1075	1163532	1164020	1163532
1076	1163995	1164294	1163995
1077	1165569	1165030	1165569
1078	1166108	1165566	1166108
1079	1166644	1166141	1166644
1080	1167055	1168374	1167055
1081	1169218	1168337	1169218
1082	1169823	1169218	1169823
1083	1171324	1170572	1171324
1084	1172085	117117	1172085
1085	1172394	1173773	1172394
1086	1175209	117388	1 1175209
1087	1175555	117512	1175360
1088	1175778	117704:	3 1175778

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ORF Nos	begin	end	potential start
1089	1177177	1179048	1177177
1090	1179156	1180085	1179156
1091	1180045	1180779	1180045
1092	1181942	1180788	1181942
1093	1182296	1181961	1182296
1094	1183844	1182300	1183844
1095	1184420	1183848	1184420
1096	1185382	1184366	1185382
1097	1185858	1185226	1185858
1098	1186164	1186481	1186185
1099	1187386	1186484	1187386
1100	1187370	1189028	1187370
1101	1189321	1190889	1189321
1102	1191142	1192146	1191142
1103	1191974	1191729	1191974
1104	1193815	1192991	1193815
1105	1195702	1194248	1195702
1106	1196303	1195716	1196303
1107	1196831	1196337	1196831
1108	1197807	1196746	1197651
1109	1198740	1197883	1198668
1110	1200232	1198721	1200232
1111	1201286	1200135	1201286
1112	1202386	1201259	1202350
1113	1202901	1202350	1202901
1114	1204162	1202816	1204162
1115	1203177	1203464	1203177
1116	1205028	1204180	1205028
1117	1206392	1204878	1206392
1118	1206742	1206086	1206742
1119	1207872	1206724	1207872
1120	1208852	1207851	1208852
112	1210518	1209742	1210518
1122	1210703	1211494	1210703

ORF Nos	begin	end	potential start
1123	1211870	1212754	1211870
1124	1212742	1214064	1212742
1125	1214046	1214858	1214046
1126	1215551	1216318	1215551
1127	1216493	1216849	1216493
1128	1217183	1219612	1217183
1129	1220068	1219673	1220068
1130	1219710	1220669	1219710
1131	1220630	1221376	1220630
1132	1221645	1223681	1221645
1133	1223894	1224988	1223900
1134	1225000	1225830	1225000
1135	1227810	1225879	1227810
1136	1226528	1226908	1226528
1137	1229972	1228311	1229972
1138	47569	47018	47569
1139	49980	49117	49980
1140	53356	52898	53356
1141	54477	54884	54477
1142	63753	63998	63753
1143	77164	77487	77164
1144	79724	79302	79724
1145	88721	88951	88721
1146	94067	94429	94067
1147	122832	123341	122832
1148	147536	147234	147536
1149	158990	159346	158990
1150	168470	168979	168470
1151	169183	169452	169204
1152	171785	171504	171785
115	172518	171775	172518
1154	193599	194045	193599
115:	195704	196075	195704
1150	210687	210145	210684
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ORF Nos	begin	end	potential start
1157	211100	210708	211100
1158	215420	215088	215420
1159	217914	218246	217914
1160	218925	218701	218925
1161	223785	223525	223785
1162	224271	223999	224271
1163	228691	228407	228691
1164	235050	235334	235050
1165	252308	253021	252308
1-166	258280	258912	258280
1167	261325	261567	261325
1168	268195	268878	268195
1169	269447	268881	269447
1170	271263	271538	271263
1171	271957	272346	271957
1172	274176	274550	274176
1173	275736	275314	275736
1174	276490	276927	276490
1175	277577	277861	277577
1176	288163	287909	288163
1177	290130	289789	290130
1178	290989	291225	290989
1179	291372	291860	291372
1180	311239	311622	311239
1181	328665	328384	328665
1182	337348	338289	337348
1183	364764	364369	364764
1184	389623	390135	389623
1185	393729	394343	393729
1186	407379	407621	407379
1187	410944	410708	410944
1188	427632	427988	427632
1189	428172	428486	428172
1190	436761	437246	436761

ORF Nos	begin	end	potential start
1191	460911	461159	460911
1192	477597	477313	477597
1193	487303	487001	487303
1194	487764	487534	487764
1195	498502	499017	498502
1196	499795	500466	499795
1197	571928	572344	571928
1198	572367	572131	572367
1199	588184	587915	588184
1200	600587	600907	600587
1201	609731	608895	609731
1202	614039	614755	614039
1203	614823	615152	614823
1204	638244	638831	638244
1205	638819	639094	638819
1206	639073	639636	639073
1207	647901	648236	647901
1208	678510	679469	678510
1209	688178	688732	688178
1210	696045	696563	696045
1211	708998	708588	708998
1212	709808	710089	709808
1213	718240	717737	718240
1214	737828	737565	737828
1215	779502	780257	779502
1216	806310	805864	806310
1217	820931	820707	820931
1218	837696	839096	837696
1219	883307	883549	883307
1220	892010	891726	892010
1221	893277	893564	893277
1222	936998	937225	936998
1223	946865	<u> </u>	1
1224	975187	975411	975187

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ORF Nos	begin	end	potential start
1225	985882	985517	985882
1226	987713	987180	987713
1227	988215	987733	988215
1228	988754	988530	988754
1229	992542	992841	992542
1230	992759	993067	992759
1231	1004247	1004528	1004268
1232	1015013	1014294	1015013
1233	1056147	1056545	1056147
1234	1077682	1078035	1077682
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1240	1124453	1124797	1124453
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1242	1164947	1164474	1164947
1243	1170457	1170053	1170457
1244	1172342	1171863	1172342
1245	1192155	1192835	1192155
1246	1192759	1192992	1192759
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1253	1228304	1228080	1228304
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1255	27609	27367	27609
1256	67206	66967	67197
1257	70612	70352	70588
1258	132703	132945	132703
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ORF Nos	begin	end	potential start
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1260	208576	208349	208576
1261	209156	208929	209156
1262	209263	209024	209263
1263	210304	210639	210304
1264	299009	299452	299030
1265	352106	351717	352061
1266	420182	419949	420170
1267	553602	553381	553602
1268	556538	556807	556538
1269	594348	593797	594342
1270	595169	594876	595160
1271	662148	662381	662160
1272	706528	706893	706528
1273	803315	803650	803339
1274	849551	849306	849551
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1283	1012924	1012694	1012924
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ORF Nos	begin	end	potential start
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Table 4

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633         2698         2699         5202         5203           634         2700         2701         5204         5205           635         2702         2703         5206         5207           636         2704         2705         5208         5209           637         2706         2707         5210         5211           638         2710         2711         5214         5215           639         2712         2713         5216         5217           640         2714         2715         5218         5219           641         2716         2717         5220         5221           642         2718         2719         5222         5223           643         2720         2721         5224         5225           644         2722         2723         5226         5227           645         2724         2725         5238         5229           646         2726         2727         5230         5231           647         2728         2729         5232         5233           649         2736         2737         5240         5241 </td <td>631</td> <td>2692</td> <td>2693</td> <td>5196</td> <td>5197</td>	631	2692	2693	5196	5197
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635         2702         2703         5206         5207           636         2704         2705         5208         5209           637         2706         2707         5210         5211           638         2710         2711         5214         5215           639         2712         2713         5216         5217           640         2714         2715         5218         5219           641         2716         2717         5220         5221           642         2718         2719         5222         5223           643         2720         2721         5224         5225           644         2722         2723         5226         5227           645         2724         2725         5228         5229           646         2726         2727         5230         5231           647         2728         2729         5232         5233           648         2732         2733         5236         5237           649         2736         2737         5240         5241           650         2738         2739         5242         5243 </td <td>633</td> <td>2698</td> <td>2699</td> <td>5202</td> <td>5203</td>	633	2698	2699	5202	5203
636         2704         2705         5208         5209           637         2706         2707         5210         5211           638         2710         2711         5214         5215           639         2712         2713         5216         5217           640         2714         2715         5218         5219           641         2716         2717         5220         5221           642         2718         2719         5222         5223           643         2720         2721         5224         5225           643         2720         2721         5224         5225           644         2722         2723         5226         5227           645         2724         2725         5228         5229           646         2726         2727         5230         5231           647         2728         2729         5232         5233           648         2732         2733         5236         5237           649         2736         2737         5240         5241           650         2738         2739         5242         5243 </td <td>634</td> <td>2700</td> <td>2701</td> <td>5204</td> <td>5205</td>	634	2700	2701	5204	5205
637         2706         2707         5210         5211           638         2710         2711         5214         5215           639         2712         2713         5216         5217           640         2714         2715         5218         5219           641         2716         2717         5220         5221           642         2718         2719         5222         5223           643         2720         2721         5224         5225           644         2722         2723         5226         5227           645         2724         2725         5228         5229           646         2726         2727         5230         5231           647         2728         2729         5232         5233           648         2732         2733         5236         5237           649         2736         2737         5240         5241           650         2738         2739         5242         5243           651         2740         2741         5244         5245           653         2742         2743         5246         5247 </td <td>635</td> <td>2702</td> <td>2703</td> <td>5206</td> <td>5207</td>	635	2702	2703	5206	5207
638         2710         2711         5214         5215           639         2712         2713         5216         5217           640         2714         2715         5218         5219           641         2716         2717         5220         5221           642         2718         2719         5222         5223           643         2720         2721         5224         5225           644         2722         2723         5226         5227           645         2724         2725         5228         5229           646         2726         2727         5230         5231           647         2728         2729         5232         5233           648         2732         2733         5236         5237           649         2736         2737         5240         5241           650         2738         2739         5242         5243           651         2740         2741         5244         5245           652         2742         2743         5246         5247           653         2744         2745         5248         5249 </td <td>636</td> <td>2704</td> <td>2705</td> <td>5208</td> <td>5209</td>	636	2704	2705	5208	5209
639         2712         2713         5216         5217           640         2714         2715         5218         5219           641         2716         2717         5220         5221           642         2718         2719         5222         5223           643         2720         2721         5224         5225           644         2722         2723         5226         5227           645         2724         2725         5228         5229           646         2726         2727         5230         5231           647         2728         2729         5232         5233           648         2732         2733         5236         5237           649         2736         2737         5240         5241           650         2738         2739         5242         5243           651         2740         2741         5244         5245           652         2742         2743         5246         5247           653         2744         2745         5248         5249           654         2748         2749         5252         5253 </td <td>637</td> <td>2706</td> <td>2707</td> <td>5210</td> <td>5211</td>	637	2706	2707	5210	5211
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1086	3674	3675	6178	6179
1087	3676	3677	6180	6181
1088	3678	3679	6182	6183
1089	3680	3681	6184	6185
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1091	3684	3685	6188	6189
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1096	3694	3695	6198	6199
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1098	3698	3699	6202	6203
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1100	3700	3701	6204	6205
1101	3704	3705	6208	6209
1102	3706	3707	6210	6211
1103	3708	3709	6212	6213
1104	3714	3715	6218	6219
1105	3720	3721	6224	6225
1106	3722	3723	6226	6227
1107	3724	3725	6228	6229
1108	3726	3727	6230	6231
1109	3728	3729	6232	6233
1110	3730	3731	6234	6235
1111	3732	3733	6236	6237
1112	3734	3735	6238	6239
1113	3736	3737	6240	6241
1114	3740	3741	6244	6245
1115	3738	3739	6242	6243
1116	3742	3743	6246	6247
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1119	3748	3749	6252	6253
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1122	3756	3757	6260	6261
1123	3758	3759	6262	6263
1124	3760	3761	6264	6265
1125	3762	3763	6266	6267
1126	3766	3767	6270	6271
1127	3770	3771	6274	6275
1128	3772	3773	6276	6277
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1177	1880	1881	4384	4385
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1181	1968	1969	4472	4473
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1199	2484	2485	4988	4989
1200	2512	2513	5016	5017
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1202	2540	2541	5044	5045
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1211	2730	2731	5234	5235
1212	2734	2735	5238	5239
1213	2746	2747	5250	5251
1214	2802	2803	5306	5307
1215	2898	2899	5402	5403
1216	2950	2951	5454	5455
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1218	3018	3019	5522	5523
1219	3098	3099	5602	5603
1220	3118	3119	5622	5623
1221	3126	3127	5630	5631
1222	3208	3209	5712	5713
1223	3242	3243	5746	5747
1224	3294	3295	5798	5799

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1234	3492	3493	5996	5997
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1258	6308	6309	6384	6385
1259	6310	6311	6386	6387
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1263	6318	6319	6394	6395
1264	6320	6321	6396	6397
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1268	6328	6329	6404	6405
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1270	6332	6333	6408	6409
1271	6334	6335	6410	6411
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1277	6346	6347	6422	6423
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1285	6362	6363	6438	6439
1286	6364	6365	6440	6441
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TABLE 5

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1303	F	2126	3023	F	838723	4743	В	462365
1304	F	5735	3024	F	841751	4744	В	461391
1305	F	3843	3025	F	839825	4745	В	463286
1306	F	7832	3026	F	842960	4746	В	461680
1307	F	5909	3027	F	841123	4747	В	463584
1308	F	8887	3028	F	843765	4748	В	462520
1309	F	7010	3029	F	841844	4749	В	464418
1310	F	10139	3030	F	844768	4750	В	463584
1311	F	8175	3031	F	842852	4751	В	465539
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1313	F	8799	3033	F	844175	4753	В	466398
1314	F	10997	3034	F	848293	4754	В	465288
1315	F	9037	3035	F	846449	4755	В	467243
1316	F	12458	3036	F	848867	4756	В	465835
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1369	F	43250	3089	F	873244	4809	В	491191
1370	F	45478	3090	F	878156	4810	В	489561
1371	F	43579	3091	<b>F</b>	876291	4811	В	491461
1372	F	46755	3092	F	879046	4812	В	490221
1373	F	44874	3093	F	877133	4813	В	492078
1374	F	47347	3094	F	880361	4814	В	490773
1375	F	45386	3095	F	878450	4815	В	492672
1376	F	47818	3096	F	882361	4816	В	491383
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1378	F	48893	3098	F	883067	4818	В	491616
1379	F	46995	3099	F	881185	4819	В	493537
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1388	F	53516	3108	F	887996	4828	В	497500
1389	F	51611	3109	F	886093	4829	В	499390
1390	F	54242	3110	F	888494	4830	В	498057
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1393	F	53159	3113	F	887201	4833	В	500508
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1403	F	57802	3123	F	890400	4843	В	506292
1404	F	61887	3124	F	892182	4844	В	504885
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1409	F	61557	3129	F	891211	4849	В	509795
1410	F	63657	3130	F	895494	4850	В	510741
1411	F	61761	3131	F	893599	4851	В	512656
1412	F	64088	3132	F	896448	4852	В	508573
1413	F	62196	3133	F	894511	4853	В	510445
1414	F	64422	3134	F	897341	4854	В	513663
1415	F	62537	3135	F	895442	4855	В	515585
1416	F	65072	3136	F	899197	4856	В	515276
1417	F	63140	3137	F	897279	4857	В	517040
1418	F	65978	3138	F	899999	4858	В	517602
1419	F	64088	3139	F	898075	4859	В	519510
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1421	F	65146	3141	F	901103	4861	В	519510
1422	F	67466	3142	F	904798	4862	В	518075
1423	F	65580	3143	F	902923	4863	В	519947
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1426	F	68609	3146	F	907564	4866	В	521416
1427	F	66688	3147	F	905665	4867	В	523319
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1435	F	71931	3155	F	909481	4875	В	528361
1436	F	76942	3156	F	912084	4876	В	527330
1437	F	75022	3157	F	910176	4877	В	529238
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1440	F	78133	3160	F	913813	4880	В	528673
1441	F	76192	3161	F	911941	4881	В	530573
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1444	F	79471	3164	F	915053	4884	В	530864
1445	F	77481	3165	F	913141	4885	В	532745
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1447	F	77816	3167	F	914731	4887	В	533776
1448	F	80236	3168	F	917500	4888	В	534199
1449	F	78356	3169	F	915594	4889	В	536103
1450	F	81108	3170	F	918615	4890	В	536674
1451	F	79182	3171	F	916715	4891	В	538552
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1464	F	86397	3184	F	923841	4904	В	542791

1465	F	84507	3185	F	921936	4905	В	544691
1466	F	88470	3186	F	924795	4906	В	543234
1467	F	86563	3187	F	922945	4907	В	545134
1468	F	89038	3188	F	925102	4908	В	543608
1469	F	87121	3189	F	923188	4909	В	545513
1470	F	91017	3190	F	926130	4910	В	546851
1471	F	89146	3191	F	924248	4911	В	548762
1472	F	93075	3192	F	927729	4912	В	549793
1473	F	91147	3193	F	925829	4913	В	551652
1474	F	93846	3194	F	928112	4914	В	547523
1475	F	91948	3195	F	926130	4915	В	549430
1476	F	94410	3196	F	929014	4916	В	550754
1477	F	92561	3197	F	927129	4917	В	552702
1478	F	95447	3198	F	930776	4918	В	551775
1479	F	93541	3199	F	928876	4919	В	553674
1480	F	96074	3200	F	931898	4920	В	552876
1481	F	94197	3201	F	929987	4921	В	554756
1482	F	97706	3202	F	932291	4922	В	555340
1483	F	95841	3203	F	930323	4923	В	557240
1484	F	98142	3204	F	933264	4924	В	555736
1485	F	96292	3205	F	931339	4925	В	557619
1486	F	99925	3206	F	935505	4926	В	558229
1487	F	98011	3207	F	933605	4927	В	560135
1488	F	101229	3208	F	936779	4928	B	558821
1489	F	99338	3209	F	934873	4929	В	560696
1490	F	101429	3210	F	937000	4930	В	559955
1491	F	99552	3211	F	935108	4931	В	561816
1492	F	102137	3212	F	938062	4932	В	561979
1493	F	100237	3213	F	936162	4933	В	563858
1494	F	102600	3214	F	938536	4934	В	561979
1495	F	100657	3215	F	936689	4935	В	563812
1496	F	103330	3216	F	938934	4936	В	564167
1497	F	101429	3217	F	937000	4937	В	566081
1498	F	103877	3218	F	939541	4938	В	565229
1499	F	101966	3219	F	937640	4939	В	567096
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4	· — ·	10/22	222	<u> </u>	040602	4040	5	ECCATO
1500	F	104336	3220	F	940603	4940	В	566419
1501	F	102469	3221	F	938681	4941	В	568318
1502	F	108182	3222	F	940758	4942	В	567974
1503	F	106280	3223	F	938826	4943	В	569872
1504	F	111814	3224	F	941387	4944	В	568753
1505	F	109911	3225	F	939470	4945	В	570655
1506	F	112412	3226	F	942261	4946	В	569707
1507	F	110553	3227	F	940373	4947	В	571605
1508	F	113442	3228	F	942563	4948	В	571285
1509	F	111571	3229	F	940654	4949	В	573207
1510	F	113891	3230	F	942807	4950	В	572080
1511	F	112010	3231	F	940907	4951	В	573948
1512	F	114990	3232	F	943510	4952	В	572628
1513	F	113112	3233	F	941608	4953	В	574524
1514	F	115684	3234	F	943771	4954	В	573563
1515	F	113776	3235	F	941872	4955	В	575436
1516	F	116526	3236	F	944330	4956	В	572628
1517	F	114656	3237	F	942413	4957	В	574524
1518	F	117731	3238	F	945147	4958	В	575279
1519	F	115825	3239	F	943262	4959	В	577202
1520	F	118292	3240	F	945527	4960	В	576190
1521	F	116389	3241	F	943620	4961	В	578039
1522	F	119593	3242	F	946627	4962	В	578174
1523	F	117685	3243	F	944741	4963	В	580011
1524	F	120231	3244	F	947165	4964	В	579148
1525	F	118292	3245	F	945278	4965	В	581040
1526	F	122278	3246	F	948674	4966	В	580227
1527	F	120382	3247	F	946774	4967	В	582047
1528	F	122610	3248	F	949646	4968	В	580656
1529	F	120682	3249	F	947716	4969	В	582542
1530	F	123309	3250	F	950731	4970	В	580420
1531	F	121390	3251	F	948837	4971	В	582322
1532	F	126113	3252	F	951418	4972	В	581322
1533	F	124213	3253	F	949545	4973	В	583212
1534	F	128975	3254	F	951940	4974	В	582051
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1535	F	127091	3255	F	950034	4975	В	583973
1536	F	134603	3256	F	952365	4976	В	582592
1537	F	132806	3257	F	950461	4977	В	584513
1538	F	136249	3258	F	953230	4978	В	583651
1539	F	134352	3259	F	951316	4979	В	585588
1540	F	137680	3260	F	954978	4980	В	584932
1541	F	135756	3261	F	953125	4981	В	586813
1542	F	137680	3262	F	955613	4982	В	585457
1543	F	135799	3263	F	953697	4983	В	587360
1544	F	138035	3264	F	956989	4984	В	587145
1545	F	136135	3265	F	955136	4985	В	589063
1546	F	139266	3266	F	957684	4986	B_	588150
1547	F	137363	3267	F	955778	4987	В	590044
1548	F	140208	3268	F	959156	4988	В	588404
1549	F	138351	3269	F	957187	4989	В	590304
1550	F	141636	3270	F	960035	4990	В	589320
1551	F	139735	3271	F	958117	4991	В	591193
1552	F	142808	3272	F	961584	4992	В	590733
1553	F	140900	3273	F	959727	4993	В	592677
1554	F	144272	3274	F	965172	4994	В	592682
1555	F	142372	3275	F	963269	4995	В	594583
1556	F	145217	3276	F	966747	4996	В	593126
1557	F	143335	3277	F	964843	4997	В	595026
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1559	F	144645	3279	F	966111	4999	В	595882
1560	F	146965	3280	F	968508	5000	В	594521
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1562	F	147455	3282	F	969289	5002	. B	596170
1563	F	145501	3283	F	967389	5003	В	598096
1564	F	148810	3284	F	969537	5004	В	596532
1565	F	146904	3285	F	967640	5005	В	598451
1566	F	151964	3286	F	970078	5000	5 B	597438
1567	F	150062	3287	F	968137	500	7 B	599365
1568	F	154064	3288	F	970317	500	8 B	598191
1569	F	152113	3289	F	968394	500	9 B	600088
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1570	F	154888	3290	F	970857	5010	В	598836
1571	F	152963	3291	F	968969	5011	В	600749
1572	F	155418	3292	F	971657	5012	В	599476
1573	F	153558	3293	F	969757	5013	В	601327
1574	F	156528	3294	F	974954	5014	В	600192
1575	F	154606	3295	F	973067	5015	В	602103
1576	F	157433	3296	F	975200	5016	В	601131
1577	F	155516	3297	F	973300	5017	В	603030
1578	F	158771	3298	F	976362	5018	В	602307
1579	F	156842	3299	F	974418	5019	В	604209
1580	F	159105	3300	F	977009	5020	В	602810
1581	F	157219	3301	F	975050	5021	В	604759
1582	F	159657	3302	F	978153	5022	В	603529
1583	F	157761	3303	F	976255	5023	В	605402
1584	F	160240	3304	F	980532	5024	В	604759
1585	F	158316	3305	F	978632	5025	В	606662
1586	F	160675	3306	F	981701	5026	В	606076
1587	F	158778	3307	F	979785	5027	В	608046
1588	F	161289	3308	F	982885	5028	В	606843
1589	F	159402	3309	F	980983	5029	В	608746
1590	F	161918	3310	F	983878	5030	В	607504
1591	F	159979	3311	F	981973	5031	В	609404
1592	F	162214	3312	F	985264	5032	В	609224
1593	F	160297	3313	F	983395	5033	В	611138
1594	F	163996	3314	F	986953	5034	В	609952
1595	F	162045	3315	F	985049	5035	В	611865
1596	F	165189	3316	F	985623	5036	В	611138
1597	F	163288	3317	F	983760	5037	В	613033
1598	F	166730	3318	F	986956	5038	В	612012
1599	F	164828	3319	F	985049	5039	В	613917
1600	F	168243	3320	F	987506	5040	В	612554
1601	F	166327	3321	F	985592	5041	В	614453
1602	F	168907	3322	F	988307	5042	В	614136
1603	F	167064	3323	F	986404	5043	В	616017
1604	F	169129	3324	F	988783	5044	В	614978
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1605	F	167294	3325	F	986927	5045	В	616936
1606	F	170632	3326	F	989593	5046	В	615399
1607	F	168692	3327	F	987694	5047	В	617342
1608	F	171229	3328	F	990733	5048	В	616565
1609	F	169381	3329	F	988783	5049	В	618402
1610	F	171553	3330	F	991559	5050	В	617618
1611	F	169614	3331	F	989675	5051	В	619515
1612	F	172433	3332	F	992323	5052	В	619027
1613	F	170533	3333	F	990421	5053	В	620937
1614	F	173217	3334	F	992522	5054	В	620142
1615	F	171316	3335	F	990640	5055	В	622052
1616	F	174567	3336	F	993308	5056	В	620230
1617	F	172680	3337	F	991361	5057	В	622124
1618	F	175342	3338	F	992795	5058	В	621498
1619	F	173479	3339	F	990919	5059	В	623385
1620	F	175709	3340	F	994573	5060	В	622583
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1622	F	176909	3342	F	995517	5062	В	623718
1623	F	175009	3343	F	993570	5063	В	625598
1624	F	176704	3344	F	996518	5064	В	624533
1625	F	174761	3345	F	994660	5065	В	626462
1626	F	177608	3346	F	997317	5066	В	625020
1627	F	175709	3347	F	995450	5067	В	626893
1628	F	179259	3348	F	998653	5068	В	625774
1629	F	177384	3349	F	996762	5069	В	627660
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1631	F	177800	3351	F	997908	5071	В	628010
1632	F	181629	3352	F	1001112	5072	В	626646
1633	F	179743	3353	F	999238	5073	В	628522
1634	F	182851	3354	F	1001651	5074	В	628020
1635	F	180952	3355	F	999731	5075	В	629982
1636	F	184230	3356	F	1003237	5076	В	628882
1637	F	182335	3357	F	1001317	5077	В	630730
1638	F	184870	3358	F	1004049	5078	В	629982
1639	F	182962	3359	F	1002132	5079	В	631822

1640	F	185241	3360	F	1004252	5080	В	631862
1641	F	183348	3361	F	1002307	5081	В	633762
1642	F	185611	3362	F	1005400	5082	В	633774
1643	F	183685	3363	F	1003518	5083	В	635675
1644	F	186336	3364	F	1005892	5084	В	637192
1645	F	184445	3365	F	1003958	5085	В	639082
1646	F	188059	3366	F	1006516	5086	В	638321
1647	F	186171	3367	F	1004599	5087	В	640221
1648	F	190828	3368	F	1007332	5088	В	639082
1649	F	188956	3369	F	1005446	5089	В	640954
1650	F	191294	3370	F	1009066	5090	В	639317
_1.65_1	F	189428	3371	F	1007190	5091	В	641243
1652	F	192686	3372	F	1014072	5092	В	639860
1653	F	190788	3373	F	1012172	5093	В	641780
1654	F	193380	3374	F	1015614	5094	В	640868
1655	F	191474	3375	F	1013733	5095	В	642770
1656	F	193388	3376	F	1016078	5096	В	641243
1657	F	191474	3377	F	1014172	5097	В	643106
1658	F	193977	3378	F	1015924	5098	В	641605
1659	F	192059	3379	F	1014059	5099	В	643503
1660	F	195480	3380	F	1016230	5100	В	642538
1661	F	193585	3381	F	1014330	5101	В	644407
1662	F	195868	3382	F	1017479	5102	В	643243
1663	F	193969	3383	F	1015558	5103	В	645145
1664	F	197913	3384	F	1018915	5104	В	643550
1665	F	196013	3385	F	1017003	5105	В	645450
1666	F	199088	3386	F	1019328	5106	В	643925
1667	F	197213	3387	F	1017440	5107	В	645837
1668	F	202776	3388	F	1020813	5108	В	645848
1669	F	200876	3389	F	1018915	5109	В	647759
1670	F	204467	3390	F	1021621	5110	В	645987
1671	F	202497	3391	F	1019671	5111	В	647969
1672	F	205584	3392	F	1023996	5112	В	646490
1673	F	203664	3393	F	1022107	5113	В	648429
	F	206940	3394	F	1024277	5114	B	646973

1675	F	205063	3395	F	1022385	5115	В	648871
1676	F	207560	3396	F	1025368	5116	В	648115
1677	F	205587	3397	F	1023468	5117	В	650007
1678	F	208048	3398	F	1026671	5118	В	648516
1679	F	206139	3399	F	1024821	5119	В	650374
1680	F	209923	3400	F	1027688	5120	В	650567
1681	F	208023	3401	F	1025823	5121	В	652472
1682	F	210455	3402	F	1030916	5122	В	651251
1683	F	208569	3403	F	1029047	5123	В	653140
1684	F	211049	3404	F	1031342	5124	В	653186
1685	F	209147	3405	F	1029430	5125	В	655076
1686	F	211596	3406	F	1032795	5126	В	653628
1687	F	209705	3407	F	1030916	5127	В	655515
1688	F	212226	3408	F	1032978	5128	В	656010
1689	F	210311	3409	F	1031078	5129	В	657870
1690	F	213832	3410	F	1033730	5130	В	656761
1691	F	211960	3411	F	1031839	5131	В	658636
1692	F	214866	3412	F	1035774	5132	В	658389
1693	F	212921	3413	F	1033821	5133	В	660295
1694	F	215173	3414	F	1036884	5134	В	660436
1695	F	213307	3415	F	1034954	5135	В	662352
1696	F	215800	3416	F	1037476	5136	В	663483
1697	F	213957	3417	F	1035577	5137	В	665358
1698	F	216489	3418	F	1037714	5138	В	664701
1699	F	214549	3419	F	1035847	5139	В	666607
1700	F	216980	3420	F	1038782	5140	В	665978
1701	F	215100	3421	F	1036884	5141	В	667856
1702	F	217665	3422	F	1040777	5142	В	667238
1703	F	215793	3423	F	1038856	5143	В	669172
1704	F	218039	3424	F	1042132	5144	В	668195
1705	F	216071	3425	F	1040216	5145	В	670046
1706	F	218476	3426	F	1043148	5146	В	668791
1707	F	216560	3427	F	1041215	5147	В	670691
1708	F	218769	3428	F	1044388	5148	В	669426
1709	F	216809	3429	F	1042445	5149	В	671326
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1711	F	218128	3431	F	1043224	5151	В	673055
1712	F	221210	3432	F	1046223	5152	В	671659
1713	F	219275	3433	F	1044324	5153	В	673547
1714	F	222497	3434	F	1047299	5154	В	672474
1715	F	220601	3435	F	1045364	5155	В	674347
1716	F	223292	3436	F	1049803	5156	В	673238
1717	F	221403	3437	F	1047914	5157	В	675140
1718	F	223775	3438	F	1050341	5158	В	674944
1719	F	221877	3439	F	1048431	5159	В	676911
1720	F	224250	3440	F	1050862	5160	В	674797
1721	F	222377	3441	F	1048907	5161	В	676669
1722	F	224906	3442	F	1051515	5162	В	675741
1723	F	223008	3443	F	1049572	5163	В	677643
1724	F	225283	3444	F	1051828	5164	В	676340
1725	F	223418	3445	F	1049917	5165	В	678204
1726	F	226670	3446	F	1052885	5166	В	676911
1727	F	224770	3447	F	1050957	5167	В	678770
1728	F	227849	3448	F	1053963	5168	В	677240
1729	F	225937	3449	F	1052057	5169	В	679136
1730	F	228185	3450	F	1055238	5170	В	677873
1731	F	226269	3451	F	1053362	5171	В	679767
1732	F	228393	3452	F	1055849	5172	В	678549
1733	F	226512	3453	F	1053963	5173	В	680420
1734	F	229334	3454	F	1056332	5174	В	679692
1735	F	227499	3455	F	1054465	5175	В	681628
1736	F	230761	3456	F	1056738	5176	В	680320
1737	F	228846	3457	F	1054830	5177	В	682220
1738	F	231287	3458	F	1058019	5178	В	681126
1739	F	229334	3459	F	1056110	5179	В	683046
1740	F	231731	3460	F	1058504	5180	В	682558
1741	F	229927	3461	F	1056587	5181	В	684404
1742	F	232865	3462	F	1059300	5182	В	681857
1743	F	231027	3463	F	1057406	5183	В	683768
1744	F	232865	3464	F	1060356	5184	В	683046
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1745	F	231027	3465	F	1058400	5185	В	684944
1746	F	234315	3466	F	1061455	5186	В	684128
1747	F	232394	3467	F	1059456	5187	В	686124
1748	F	234823	3468	F	1062092	5188	В	684893
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1750	F	235154	3470	F	1063884	5190	В	685389
1751	F	233245	3471	F	1061983	5191	В	687290
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1753	F	234520	3473	F	1063056	5193	В	688106
1754	F	237268	3474	F	1067125	5194	В	687534
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1756	F	238047	3476	F	1067963	5196	В	688416
1757	F	236162	3477	F	1066075	5197	В	690275
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1759	F	236736	3479	F	1066668	5199	В	690855
1760	F	239957	3480	F	1069752	5200	В	689727
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1762	F	241373	3482	F	1071068	5202	В	690496
1763	F	239482	3483	F	1069210	5203	В	692386
1764	F	242017	3484	F	1072701	5204	В	691349
1765	F	240072	3485	F	1070806	5205	В	693249
1766	F	242740	3486	F	1073987	5206	В	692864
1767	F	240829	3487	F	1072090	5207	В	694724
1768	F	243281	3488	F	1075643	5208	В	695287
1769	F	241373	3489	F	1073742	5209	В	697187
1770	F	244244	3490	F	1076350	5210	В	696275
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1773	F	244179	3493	F	1075555	5213	В	698696
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1775	F	245697	3495	F	1075880	5215	В	700090
1776	F	249216	3496	F	1078445	5216	В	700037
1777	F	247244	3497	F	1076529	5217	В	701923
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1779	F	249137	3499	F	1077523	5219	В	704050

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1780	F	252064	3500	F	1079715	5220	В	703443
1781	F	250189	3501	F	1077850	5221	В	705316
1782	F	252900	3502	F	1080538	5222	В	704441
1783	F	251000	3503	F	1078655	5223	В	706351
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1785	F	251855	3505	F	1079228	5225	В	707413
1786	F	254993	3506	F	1083006	5226	В	706312
1787	F	253138	3507	F	1081108	5227	В	708190
1788	F	256414	3508	F	1084404	5228	В	707058
1789	F	254509	3509	F	1082465	5229	В	708979
1790	F	257283	3510	F	1085896	5230	В	707856
1791	F	255383	3511	F	1083990	5231	В	709719
1792	F	257279	3512	F	1086468	5232	В	708906
1793	F	255379	3513	F	1084563	5233	В	710811
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1795	F	256107	3515	F	1085985	5235	В	711132
1796	F	259005	3516	F	1088427	5236	В	710074
1797	F	257128	3517	F	1086527	5237	В	711924
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1800	F	261551	3520	F	1089668	5240	В	711748
1801	F	259650	3521	F	1087768	5241	В	713690
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1803	F	260611	3523	F	1090767	5243	В	714407
1804	F	262960	3524	F	1093357	5244	В	715001
1805	F	261060	3525	F	1091465	5245	В	716854
1806	F	264509	3526	F	1093957	5246	В	715983
1807	F	262614	3527	F	1092070	5247	В	717887
1808	F	265837	3528	F	1095818	5248	В	717800
1809	F	263925	3529	F	1093955	5249	В	719668
1810	F	266239	3530	F	1096359	5250	В	718468
1811	F	264367	3531	F	1094509	5251	В	720383
1812	F	267185	3532	F	1097047	5252	В	720469
1813	F	265286	3533	F	1095114	5253	В	722367
1814	F	267909	3534	F	1097365	5254	В	722645
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1817	F	266756	3537	F	1095767	5257	В	725273
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1819	F	267505	3539	F	1096242	5259	В	725691
1820	F	271044	3540	F	1098560	5260	В	724469
1821	F	269121	3541	F	1096663	5261	В	726387
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1825	F	270645	3545	F	1097547	5265	В	727988
1826	F	273007	3546	F	1100878	5266	В	727397
1827	F	271098	3547	F	1098942	5267	В	729236
1828	F	273463	3548	F	1101839	5268	В	728347
1829	F	271500	3549	F	1099956	5269	В	730278
1830	F	273922	3550	F	1104621	5270	В	728816
1831	F	272057	3551	F	1102789	5271	В	730718
1832	F	275083	3552	F	1106487	5272	В	729846
1833	F	273094	3553	F	1104562	5273	В	731740
1834	F	275495	3554	F	1107225	5274	В	730005
1835	F	273554	3555	F	1105318	5275	В	731898
1836	F	275739	3556	F	1107814	5276	В	730377
1837	F	273878	3557	F	1105922	5277	В	732272
1838	F	276229	3558	F	1108282	5278	В	730759
1839	F	274371	3559	F	1106374	5279	В	732659
1840	F	276548	3560	F	1113162	5280	В	732249
1841	F	274638	3561	F	1111308	5281	В	734124
1842	F	277098	3562	F	1114813	5282	В	732647
1843	F	275178	3563	F	1112949	5283	В	734590
1844	F	277358	3564	F	1116611	5284	В	733144
1845	F	275448	3565	F	1114766	5285	В	735088
1846	F	277609	3566	F	1118605	5286	В	733858
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1848	F	278314	3568	F	1119754	5288	В	734124
1849	F	276386	3569	F	1117874	5289	В	736028
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1850	F	279310	3570	F	1120291	5290	В	734523
1851	F	277385	3571	F	1118385	5291	В	736441
1852	F	280627	3572	F	1121099	5292	В	735088
1853	F	278702	3573	F	1119202	5293	В	736978
1854	F	281471	3574	F	1121886	5294	В	735416
1855	F	279559	3575	F	1119982	5295	В	737315
1856	F	282239	3576	F	1122979	5296	В	735822
1857	F	280288	3577	F	1121038	5297	В	737700
1858	F	283832	3578	F	1123376	5298	В	736099
1859	F	281933	3579	F	1121486	5299	В	737981
1860	F	284384	3580	F	1124136	5300	В	736714
1861	F	282486	3581	F	1122333	5301	В	738612
1862	F	285373	3582	F	1124623	5302	В	737448
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1864	F	285919	3584	F	1125306	5304	В	737802
1865	F	284059	3585	F	1123423	5305	В	739693
1866	F	286742	3586	F	1126300	5306	В	738048
1867	F	284879	3587	F	1124399	5307	В	739948
1868	F	287216	3588	F	1127440	5308	В	738964
1869	F	285329	3589	F	1125545	5309	В	740808
1870	F	287671	3590	F	1128968	5310	В	739282
1871	F	285751	3591	F	1127134	5311	В	741190
1872	F	288273	3592	F	1129916	5312	В	739956
1873	F	286323	3593	F	1128111	5313	В	741906
1874	F	288618	3594	F	1131255	5314	В	740743
1875	F	286685	3595	F	1129330	5315	В	742597
1876	F	288273	3596	F	1132598	5316	В	741190
1877	F	286323	3597	F	1130684	5317	В	743081
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1879	F	287836	3599	F	1132002	5319	В	743875
1880	F	289508	3600	F	1134373	5320	В	743009
1881	F	287667	3601	F	1132510	5321	В	744914
1882	F	290750	3602	F	1135431	5322	В	743875
1883	F	288858	3603	F	1133531	5323	В	745738
1884	F	291142	3604	F	1135730	5324	В	744325
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1886       F       291702       3606       F       1136932       5326       H         1887       F       289812       3607       F       1135040       5327       H         1888       F       292522       3608       F       1139875       5328       H         1889       F       290633       3609       F       1137942       5329       H         1890       F       293035       3610       F       1141133       5330       H         1891       F       291142       3611       F       1139231       5331       H         1892       F       293731       3612       F       1142301       5332       H         1893       F       291786       3613       F       1140366       5333       H         1894       F       294530       3614       F       1143505       5335       H         1895       F       292670       3615       F       1146637       5336       H         1897       F       292513       3618       F       1147417       5338       H         1898       F       296092       3618       F	B B B B B B B B B B B B B B B B B B B	746234 744824 746724 745207 747073 746828 748738 747344 749206 748253 750094 748856 750717
1887       F       289812       3607       F       1135040       5327       H         1888       F       292522       3608       F       1139875       5328       H         1889       F       290633       3609       F       1137942       5329       H         1890       F       293035       3610       F       1141133       5330       H         1891       F       291142       3611       F       1139231       5331       H         1892       F       293731       3612       F       1142301       5332       H         1893       F       291786       3613       F       1140366       5333       H         1894       F       294530       3614       F       1145346       5334       H         1895       F       292670       3615       F       1143505       5335       H         1896       F       294367       3616       F       1146637       5336       H         1897       F       292513       3618       F       1147417       5338       H         1898       F       296092       3618       F	B B B B B B B B B B B B B B B B B B B	746724 745207 747073 746828 748738 747344 749206 748253 750094 748856
1888       F       292522       3608       F       1139875       5328       H         1889       F       290633       3609       F       1137942       5329       H         1890       F       293035       3610       F       1141133       5330       H         1891       F       291142       3611       F       1139231       5331       H         1892       F       293731       3612       F       1142301       5332       H         1893       F       291786       3613       F       1140366       5333       H         1894       F       294530       3614       F       1145346       5334       H         1895       F       292670       3615       F       1143505       5335       H         1896       F       294367       3616       F       1146637       5336       H         1897       F       292513       3618       F       1147417       5338       H         1898       F       296092       3618       F       1147417       5338       H	B B B B B B B B B B B B B B B B B B B	745207 747073 746828 748738 747344 749206 748253 750094 748856
1889       F       290633       3609       F       1137942       5329       I         1890       F       293035       3610       F       1141133       5330       I         1891       F       291142       3611       F       1139231       5331       I         1892       F       293731       3612       F       1142301       5332       I         1893       F       291786       3613       F       1140366       5333       I         1894       F       294530       3614       F       1145346       5334       I         1895       F       292670       3615       F       1143505       5335       I         1896       F       294367       3616       F       1146637       5336       I         1897       F       292513       3617       F       1144743       5337       I         1898       F       296092       3618       F       1147417       5338       I	B B B B B B B B B	747073 746828 748738 747344 749206 748253 750094 748856
1890       F       293035       3610       F       1141133       5330       1         1891       F       291142       3611       F       1139231       5331       1         1892       F       293731       3612       F       1142301       5332       1         1893       F       291786       3613       F       1140366       5333       1         1894       F       294530       3614       F       1145346       5334       1         1895       F       292670       3615       F       1143505       5335       1         1896       F       294367       3616       F       1146637       5336       1         1897       F       292513       3617       F       1144743       5337       1         1898       F       296092       3618       F       1147417       5338       1	B B B B B B B B	746828 748738 747344 749206 748253 750094 748856
1891       F       291142       3611       F       1139231       5331       I         1892       F       293731       3612       F       1142301       5332       I         1893       F       291786       3613       F       1140366       5333       I         1894       F       294530       3614       F       1145346       5334       I         1895       F       292670       3615       F       1143505       5335       I         1896       F       294367       3616       F       1146637       5336       I         1897       F       292513       3617       F       1144743       5337       I         1898       F       296092       3618       F       1147417       5338       I	B B B B B B B	748738 747344 749206 748253 750094 748856
1892       F       293731       3612       F       1142301       5332       1         1893       F       291786       3613       F       1140366       5333       1         1894       F       294530       3614       F       1145346       5334       1         1895       F       292670       3615       F       1143505       5335       1         1896       F       294367       3616       F       1146637       5336       1         1897       F       292513       3617       F       1144743       5337       1         1898       F       296092       3618       F       1147417       5338       1	B B B B B B	747344 749206 748253 750094 748856
1893       F       291786       3613       F       1140366       5333       1         1894       F       294530       3614       F       1145346       5334       1         1895       F       292670       3615       F       1143505       5335       1         1896       F       294367       3616       F       1146637       5336       1         1897       F       292513       3617       F       1144743       5337       1         1898       F       296092       3618       F       1147417       5338       1	B B B B B	749206 748253 750094 748856
1894     F     294530     3614     F     1145346     5334     1       1895     F     292670     3615     F     1143505     5335     1       1896     F     294367     3616     F     1146637     5336     1       1897     F     292513     3617     F     1144743     5337     1       1898     F     296092     3618     F     1147417     5338     1	B B B B	748253 750094 748856
1895     F     292670     3615     F     1143505     5335     1       1896     F     294367     3616     F     1146637     5336     1       1897     F     292513     3617     F     1144743     5337     1       1898     F     296092     3618     F     1147417     5338     1	B B B	750094 748856
1896     F     294367     3616     F     1146637     5336     1       1897     F     292513     3617     F     1144743     5337     1       1898     F     296092     3618     F     1147417     5338     1	B B	748856
1897     F     292513     3617     F     1144743     5337     1       1898     F     296092     3618     F     1147417     5338     1	В	
1898 F 296092 3618 F 1147417 5338		750717
	R	/30/1/
1899 F 294209 3619 F 1145547 5339	ا ت	749376
1000	В	751265
1900 F 297611 3620 F 1147981 5340	В	750180
1901 F 295757 3621 F 1146086 5341	В	752086
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1903 F 296092 3623 F 1146211 5343	В	752569
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1907 F 297511 3627 F 1147890 5347	В	755162
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1917 F 301730 3637 F 1153548 5357	В	763356
1918 F 304350 3638 F 1156630 5358	В	762077
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1921	F	303226	3641	F	1155862	5361	В	764410
1922	F	306244	3642	F	1160695	5362	В	763118
1923	F	304350	3643	F	1158788	5363	В	765018
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1925	F	305310	3645	F	1160468	5365	В	765504
1926	F	307799	3646	F	1163300	5366	В	764000
1927	F	305877	3647	F	1161413	5367	В	765907
1928	F	309173	3648	F	1163763	5368	В	765391
1929	F	307301	3649	F	1161842	5369	В	767328
1930	F	310158	3650	F	1164224	5370	В	767041
1931	F	308306	3651	F	1162283	5371	В	768951
1932	F	311020	3652	F	1164800	5372	В	768271
1933	F	309118	3653	F	1162908	5373	В	770171
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1935	F	309126	3655	F	1163427	5375	В	770686
1936	F	311552	3656	F	1165877	5376	В	769562
1937	F	309658	3657	F	1163960	5377	В	771608
1938	F	312510	3658	F	1166827	5378	В	770752
1939	F	310614	3659	F	1164936	5379	В	772652
1940	F	313134	3660	F	1168099	5380	В	771701
1941	F	311255	3661	F	1166212	5381	В	773620
1942	F	313674	3662	F	1168991	5382	В	773316
1943	F	311717	3663	F	1167093	5383	В	775178
1944	F	314490	3664	F	1169769	5384	В	773690
1945	F	312633	3665	F	1167907	5385	В	775579
1946	F	315306	3666	F	1170349	5386	В	774596
1947	F	313355	3667	F	1168446	5387	В	776522
1948	F	315932	3668	F	1170953	5388	В	776300
1949	F	314033	3669	F	1169031	5389	В	778224
1950	F	318434	3670	F	1171641	5390	В	775346
1951	F	316516	3671	F	1169703	5391	В	777266
1952	F	320876	3672	F	1172172	5392	В	775618
1953	F	318949	3673	F	1170256	5393	В	777518
1954	F	321403	3674	F	1173649	5394	В	777266
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1960	F	323634	3680	F	1176927	5400	В	779557
1961	F	321726	3681	F	1175025	5401	В	781481
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1963	F	323211	3683	F	1176985	5403	В	782380
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1966	F.	327607	3686	F	1180498	5406	В	781747
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1969	F	326262	3689	F	1179839	5409	В	784912
1970	F	328630	3690	F	1182069	5410	В	783820
1971	F	326723	3691	F	1180140	5411	В	785752
1972	F	329134	3692	F	1183626	5412	В	785255
1973	F	327178	3693	F	1181716	5413	В	787155
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1977	F	330252	3697	F	1183084	5417	В	790554
1978	F	334575	3698	F	1185897	5418	В	789164
1979	F	332660	3699	F	1184029	5419	В	791064
1980	F	335884	3700	F	1187151	5420	В	790001
1981	F	333980	3701	F	1185251	5421	В	791900
1982	F	337129	3702	F	1186262	5422	В	791734
1983	F	335202	3703	F	1184361	5423	В	793679
1984	F	337910	3704	F	1189054	5424	В	792944
1985	F	335955	3705	F	1187160	5425	В	794875
1986	F	338746	3706	F	1190885	5426	В	793809
1987	F	336795	3707	F	1188990	5427	В	795692
1988	F	339217	3708	F	1191507	5428	В	794580
1989	F	337362	3709	F	1189579	5429	В	796450
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1991   F   338083   3711   F   1190008   5431   B   796966   1992   F   343144   3712   F   1192524   5432   B   795956   1993   F   341266   3713   F   1190640   5433   B   797855   1994   F   343699   3714   F   1192759   5434   B   797018   1995   F   341813   3715   F   1190869   5435   B   798918   1996   F   344108   3716   F   1193642   5436   B   798989   1997   F   342204   3717   F   1191742   5437   B   800875   1998   F   344851   3718   F   1193557   5438   B   800069   1999   F   342933   3719   F   1191657   5439   B   801944   2000   F   346483   3720   F   1194015   5440   B   799840   2001   F   344219   3721   F   1192120   5441   B   801701   2002   F   346493   3723   F   1193560   5443   B   803445   2004   F   346815   3724   F   1196093   5444   B   802717   2005   F   344907   3725   F   119415   5446   B   803559   2007   F   345956   3727   F   1194592   5447   B   805419   2008   F   350856   3730   F   1195724   5449   B   805419   2010   F   35856   3730   F   1198499   5450   B   805383   2011   F   348951   3731   F   1195724   5449   B   805333   2011   F   348951   3731   F   1196578   5452   B   806107   2012   F   352008   3732   F   119912   5452   B   806107   2015   F   355320   3734   F   1200699   5454   B   807291   2015   F   355321   3735   F   119913   5455   B   808430   2016   F   355223   3740   F   1200694   5460   B   808533   2016   F   355232   3740   F   1200694   5461   B   807291   2022   F   355331   3742   F   1200694   5461   B   807291   2022   F   355393   3742   F   1200694   5461   B   807291   2022   F   355393   3742   F   1200694   5463   B   808972   2024   F   355391   3744   F   1200694   5464   B   809674	1990	F	339999	3710	F	1191932	5430	В	795066
1993         F         341266         3713         F         119040         5433         B         797855           1994         F         343699         3714         F         1192759         5434         B         797018           1995         F         341813         3715         F         1190869         5435         B         798918           1996         F         344108         3716         F         1193642         5436         B         798989           1997         F         342204         3717         F         1191742         5437         B         800875           1998         F         344851         3718         F         1193557         5438         B         800069           1999         F         342933         3719         F         1196157         5439         B         801944           2000         F         346148         3720         F         1194015         5440         B         799840           2001         F         346493         3722         F         1192120         5441         B         801701           2002         F         346815         3724         F<		F			F	1190008	5431	В	796966
1994         F         343699         3714         F         1192759         5434         B         797018           1995         F         341813         3715         F         1190869         5435         B         798918           1996         F         344108         3716         F         1193642         5436         B         798989           1997         F         342204         3717         F         1191742         5437         B         800875           1998         F         344851         3718         F         1193557         5438         B         800069           1999         F         342933         3719         F         1191657         5439         B         801944           2000         F         346483         3720         F         1194015         5440         B         799840           2001         F         346493         3722         F         1195490         5442         B         801533           2003         F         344590         3723         F         1195690         5443         B         803445           2004         F         346815         3724         F	1992	F	343144	3712	F	1192524	5432	В	795956
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1996         F         344108         3716         F         1193642         5436         B         798989           1997         F         342204         3717         F         1191742         5437         B         800875           1998         F         344851         3718         F         1193557         5438         B         800069           1999         F         342933         3719         F         1191657         5439         B         801944           2000         F         346148         3720         F         1194015         5440         B         799840           2001         F         346493         3722         F         1195490         5441         B         801701           2002         F         346493         3722         F         1195490         5442         B         801533           2004         F         346815         3724         F         1196093         5443         B         803445           2005         F         344907         3725         F         1194215         5445         B         804581           2006         F         347836         3727         F	1994	F	343699	3714	F	1192759	5434	В	797018
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1998         F         344851         3718         F         1193557         5438         B         800069           1999         F         342933         3719         F         1191657         5439         B         801944           2000         F         346148         3720         F         1194015         5440         B         799840           2001         F         344219         3721         F         1192120         5441         B         801701           2002         F         346493         3722         F         1195490         5442         B         801533           2003         F         344590         3723         F         1193560         5443         B         803445           2004         F         346815         3724         F         1196093         5444         B         802717           2005         F         344907         3725         F         1194215         5445         B         804581           2006         F         347836         3726         F         1194592         5447         B         805419           2007         F         348432         3729         F	1996	F	344108	3716	F	1193642	5436	В	798989
1999	1997	F	342204	3717	F	1191742	5437	В	800875
2000         F         346148         3720         F         1194015         5440         B         799840           2001         F         344219         3721         F         1192120         5441         B         801701           2002         F         346493         3722         F         1195490         5442         B         801533           2003         F         344590         3723         F         1193560         5443         B         803445           2004         F         346815         3724         F         1196093         5444         B         802717           2005         F         344907         3725         F         1194215         5445         B         804581           2006         F         347836         3726         F         1196474         5446         B         803559           2007         F         345956         3727         F         1194592         5447         B         805419           2008         F         350379         3728         F         1197659         5448         B         804032           2009         F         348432         3729         F	1998	F	344851	3718	F	1193557	5438	В	800069
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2002         F         346493         3722         F         1195490         5442         B         801533           2003         F         344590         3723         F         1193560         5443         B         803445           2004         F         346815         3724         F         1196093         5444         B         802717           2005         F         344907         3725         F         1194215         5445         B         804581           2006         F         347836         3726         F         1196474         5446         B         803559           2007         F         345956         3727         F         1194592         5447         B         805419           2008         F         350379         3728         F         1197659         5448         B         804032           2009         F         348432         3729         F         1195724         5449         B         805931           2010         F         350856         3730         F         1198499         5450         B         805383           2011         F         352008         3732         F	2000	F	346148	3720	F	1194015	5440	В	799840
2003         F         344590         3723         F         1193560         5443         B         803445           2004         F         346815         3724         F         1196093         5444         B         802717           2005         F         344907         3725         F         1194215         5445         B         804581           2006         F         347836         3726         F         1196474         5446         B         803559           2007         F         345956         3727         F         1194592         5447         B         805419           2008         F         350379         3728         F         1197659         5448         B         804032           2009         F         348432         3729         F         1195724         5449         B         805931           2010         F         350856         3730         F         1198499         5450         B         805383           2011         F         352008         3732         F         1199912         5452         B         806107           2013         F         350106         3733         F	2001	F	344219	3721	F	1192120	5441	В	801701
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2008         F         350379         3728         F         1197659         5448         B         804032           2009         F         348432         3729         F         1195724         5449         B         805931           2010         F         350856         3730         F         1198499         5450         B         805383           2011         F         348951         3731         F         1196578         5451         B         807291           2012         F         352008         3732         F         1199912         5452         B         806107           2013         F         350106         3733         F         1197986         5453         B         807988           2014         F         353209         3734         F         1200969         5454         B         806533           2015         F         351305         3735         F         1199133         5455         B         808430           2016         F         354224         3736         F         1202121         5456         B         80653           2017         F         352312         3737         F<	2006	F	347836	3726	F	1196474	5446	В	803559
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2011         F         348951         3731         F         1196578         5451         B         807291           2012         F         352008         3732         F         1199912         5452         B         806107           2013         F         350106         3733         F         1197986         5453         B         807988           2014         F         353209         3734         F         1200969         5454         B         806533           2015         F         351305         3735         F         1199133         5455         B         808430           2016         F         354224         3736         F         1202121         5456         B         806954           2017         F         352312         3737         F         1200227         5457         B         808724           2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F	2009	F	348432	3729	F	1195724	5449	В	805931
2012         F         352008         3732         F         1199912         5452         B         806107           2013         F         350106         3733         F         1197986         5453         B         807988           2014         F         353209         3734         F         1200969         5454         B         806533           2015         F         351305         3735         F         1199133         5455         B         808430           2016         F         354224         3736         F         1202121         5456         B         806954           2017         F         352312         3737         F         1200227         5457         B         808724           2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F         1202590         5460         B         808442           2021         F         355393         3742         F	2010	F	350856	3730	F	1198499	5450	В	805383
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2014         F         353209         3734         F         1200969         5454         B         806533           2015         F         351305         3735         F         1199133         5455         B         808430           2016         F         354224         3736         F         1202121         5456         B         806954           2017         F         352312         3737         F         1200227         5457         B         808724           2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F         1202590         5460         B         808442           2021         F         353261         3741         F         1203923         5462         B         808972           2022         F         353519         3743         F         1202049         5463         B         810896	2012	F	352008	3732	F	1199912	5452	В	806107
2015         F         351305         3735         F         1199133         5455         B         808430           2016         F         354224         3736         F         1202121         5456         B         806954           2017         F         352312         3737         F         1200227         5457         B         808724           2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F         1202590         5460         B         808442           2021         F         353261         3741         F         1200694         5461         B         810357           2022         F         355393         3742         F         1203923         5462         B         808972           2023         F         353519         3743         F         1202049         5463         B         810896	2013	F	350106	3733	F	1197986	5453	В	807988
2016         F         354224         3736         F         1202121         5456         B         806954           2017         F         352312         3737         F         1200227         5457         B         808724           2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F         1202590         5460         B         808442           2021         F         353261         3741         F         1200694         5461         B         810357           2022         F         355393         3742         F         1203923         5462         B         808972           2023         F         353519         3743         F         1202049         5463         B         810896	2014	F	353209	3734	F	1200969	5454	В	806533
2017         F         352312         3737         F         1200227         5457         B         808724           2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F         1202590         5460         B         808442           2021         F         353261         3741         F         1200694         5461         B         810357           2022         F         355393         3742         F         1203923         5462         B         808972           2023         F         353519         3743         F         1202049         5463         B         810896	2015	F	351305	3735	F	1199133	5455	В	808430
2018         F         354781         3738         F         1202957         5458         B         807133           2019         F         352871         3739         F         1201058         5459         B         809033           2020         F         355223         3740         F         1202590         5460         B         808442           2021         F         353261         3741         F         1200694         5461         B         810357           2022         F         355393         3742         F         1203923         5462         B         808972           2023         F         353519         3743         F         1202049         5463         B         810896	2016	F	354224	3736	F	1202121	5456	В	806954
2019       F       352871       3739       F       1201058       5459       B       809033         2020       F       355223       3740       F       1202590       5460       B       808442         2021       F       353261       3741       F       1200694       5461       B       810357         2022       F       355393       3742       F       1203923       5462       B       808972         2023       F       353519       3743       F       1202049       5463       B       810896	2017	F	352312	3737	F	1200227	5457	В	808724
2020       F       355223       3740       F       1202590       5460       B       808442         2021       F       353261       3741       F       1200694       5461       B       810357         2022       F       355393       3742       F       1203923       5462       B       808972         2023       F       353519       3743       F       1202049       5463       B       810896	2018	F	354781	3738	F	1202957	5458	В	807133
2021     F     353261     3741     F     1200694     5461     B     810357       2022     F     355393     3742     F     1203923     5462     B     808972       2023     F     353519     3743     F     1202049     5463     B     810896	2019	F	352871	3739	F	1201058	5459	В	809033
2022         F         355393         3742         F         1203923         5462         B         808972           2023         F         353519         3743         F         1202049         5463         B         810896	2020	F	355223	3740	F	1202590	5460	В	808442
2023 F 353519 3743 F 1202049 5463 B 810896	2021	F	353261	3741	F	1200694	5461	В	810357
	2022	F	355393	3742	F	1203923	5462	В	808972
2024 F 358901 3744 F 1204631 5464 B 809674	2023	F	353519	3743	F	1202049	5463	В	810896
	2024	F	358901	3744	F	1204631	5464	В	809674

2025	F	357001	3745	F	1202753	5465	В	811557
2026	F	356594	3746	F	1205864	5466	В	810192
2027	F	354692	3747	F	1203964	5467	В	812105
2028	F	359240	3748	F	1206483	5468	В	811472
2029	F	357374	3749	F	1204592	5469	В	813357
2030	F	359721	3750	F	1207629	5470	В	813325
2031	F	357763	3751	F	1205727	5471	В	815179
2032	F	361071	3752	F	1208802	5472	В	813133
2033	F	359240	3753	F	1206909	5473	В	815134
2034	F	363605	3754	F	1209500	5474	В	813808
2035	F	361731	3755	F	1207557	5475	В	815737
2036	F	364142	3756	F	1210483	5476	Β	815246
2037	F	362246	3757	F	1208584	5477	В	817168
2038	F	364567	3758	F	1211618	5478	В	815995
2039	F	362708	3759	F	1209745	5479	В	817892
2040	F	365039	3760	F	1212523	5480	В	817264
2041	F	363184	3761	F	1210554	5481	В	819164
2042	F	365445	3762	F	1213827	5482	В	817579
2043	F	363517	3763	F	1211927	5483	В	819491
2044	F	367040	3764	F	1214875	5484	В	818890
2045	F	365144	3765	F	1212992	5485	В	820733
2046	F	368825	3766	F	1215293	5486	В	819332
2047	F	366993	3767	F	1213430	5487	В	821217
2048	F	369698	3768	F	1216043	5488	В	820096
2049	F	367760	3769	F	1214183	5489	В	821951
2050	F	370141	3770	F	1216226	5490	В	820945
2051	F	368239	3771	F	1214374	5491	В	822870
2052	F	372329	3772	F	1216927	5492	В	821151
2053	F	370375	3773	F	1215064	5493	В	823079
2054	F	372779	3774	F	1219490	5494	В	822558
2055	F	370881	3775	F	1217534	5495	В	824449
2056	F	373223	3776	F	1219431	5496	В	823767
2057	F	371342	3777	F	1217534	5497	В	825634
2058	F	373939	3778	F	1220403	5498	В	825876
2059	F	372017	3779	F	1218475	5499	В	827737
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2061	F	372953	3781	F	1219499	5501	В	828435
2062	F	375351	3782	F	1223653	5502	В	827511
2063	F	373487	3783	F	1221767	5503	В	829428
2064	F	376316	3784	F	1224758	5504	В	828829
2065	F	374416	3785	F	1222881	5505	В	830729
2066	F	377737	3786	F	1226308	5506	В	830262
2067	F	375828	3787	F	1224409	5507	В	832158
2068	F	379537	3788	F	1225625	5508	В	831286
2069	F	377660	3789	F	1223654	5509	В	833182
2070	F	380033	3790	F	1227566	5510	В	831946
2071	F.	378160	. 3791	F	1225677	5511	В	833848
2072	F	380789	3792	F	1227858	5512	В	833372
2073	F	378889	3793	F	1225937	5513	В	835267
2074	F	381238	3794	F	1228081	5514	В	834125
2075	F	379279	3795	F	1226189	5515	В	835992
2076	F	382969	3796	В	1019	5516	В	835267
2077	F	381124	3797	В	2954	5517	В	837193
2078	F	383293	3798	В	1843	5518	В	836111
2079	F	381425	3799	В	3739	5519	В	837952
2080	F	385178	3800	В	2694	5520	В	837844
2081	F	383278	3801	В	4545	5521	В	839751
2082	F	386271	3802	В	3694	5522	В	839381
2083	F	384392	3803	В	5513	5523	В	841221
2084	F	386780	3804	В	4290	5524	В	841127
2085	F	384891	3805	В	6238	5525	В	843073
2086	F	389383	3806	В	5924	5526	В	842409
2087	F	387504	3807	В	7846	5527	В	844323
2088	F	389901	3808	В	7687	5528	В	843691
2089	F	388001	3809	В	9583	5529	В	845602
2090	F	390700	3810	В	9189	5530	В	844244
2091	F	388732	3811	В	11095	5531	В	846153
2092	F	391612	3812	В	10261	5532	В	845319
2093	F	389763	3813	В	12119	5533	В	847139
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2096	F	392540	3816	В	11463	5536	В	848760
2097	F	390639	3817	В	13355	5537	В	850653
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2099	F	391609	3819	В	14850	5539	В	851174
2100	F	393904	3820	В	14425	5540	В	850753
2101	F	392025	3821	В	16332	5541	В	852649
2102	F	394703	3822	В	17477	5542	В	851795
2103	F	392782	3823	В	19400	5543	В	853690
2104	F	395024	3824	В	16296	5544	В	852696
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2106	F	395705	3826	В	21128	5546	В	853938
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2108	F	397607	3828	В	22265	5548	В	855338
2109	F	395705	3829	В	24185	5549	В	857240
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2112	F	399848	3832	В	26350	5552	В	856786
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2115	F	399008	3835	В	28258	5555	В	860735
2116	F	401183	3836	В	27241	5556	В	859824
2117	F	399301	3837	В	29113	5557	В	861787
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2121	F	401527	3841	В	30700	5561	В	863252
2122	F	404124	3842	В	29727	5562	В	861677
2123	F	402206	3843	В	31642	5563	В	863558
2124	F	405765	3844	В	30253	5564	В	863171
2125	F	403865	3845	В	32158	5565	В	865099
2126	F	407131	3846	В	31775	5566	В	865021
2127	F	405243	3847	В	33657	5567	В	866922
2128	F	407456	3848	В	32511	5568	В	865497
2129	F	405563	3849	В	34422	5569	В	867408
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2130	F	408841	٢	3850	В	34214	Γ	5570	В	866808
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2139	F	412268	ŀ	3859	В	41496		5579	В	871880
2140	F	415013	1	3860	В	40942	1	5580	В	870857
2141	F	413111		3861	В	42840	-	5581	В	872753
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2144	F	417033		3864	В	43329		5584	В	872758
2145	F	415114		3865	В	45196	T	5585	В	874658
2146	F	417163		3866	В	44025		5586	В	874131
2147	F	415332		3867	В	45979	ſ	5587	В	876122
2148	F	418166		3868	В	45048		5588	В	874903
2149	F	416265		3869	В	46970		5589	В	876793
2150	F	420186		3870	В	45582		5590	В	875548
2151	F	418259		3871	В	47472		5591	В	877437
2152	F	420697		3872	В	45979	ſ	5592	В	878078
2153	F	418861		3873	В	47901		5593	В	880011
2154	F	421313	1	3874	В	47216		5594	В	879478
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2156	F	422172	1	3876	В	47791		5596	В	880874
2157	F	420342		3877	В	49689		5597	В	882771
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2159	F	421412	1	3879	В	50126		5599	В	884644
2160	F	424008	1	3880	В	49180		5600	В	883542
2161	F	422073		3881	В	51105	] [	5601	В	885447
2162	F	424585		3882	В	50231		5602	В	883777
2163	F	422711		3883	В	52149		5603	В	885689
2164	F	426021		3884	В	51697		5604	В	884430

2166         F         427407         3886         B         52917         5606         B         885           2167         F         425513         3887         B         54735         5607         B         887           2168         F         427936         3888         B         53619         5608         B         887           2170         F         426053         3889         B         55476         5609         B         889           2170         F         428592         3890         B         53910         5610         B         888           2171         F         426717         3891         B         55816         5611         B         889           2172         F         430475         3892         B         54416         5612         B         888           2173         F         428558         3893         B         56326         5613         B         890           2174         F         431378         3894         B         55107         5614         B         889           2175         F         429417         3895         B         56093         5616									
2167         F         425513         3887         B         54735         5607         B         887           2168         F         427936         3888         B         53619         5608         B         887           2169         F         426053         3889         B         55476         5609         B         889           2170         F         428592         3890         B         53910         5610         B         888           2171         F         426717         3891         B         55816         5611         B         890           2173         F         428558         3893         B         56326         5613         B         890           2174         F         431378         3894         B         55107         5614         B         889           2175         F         429417         3895         B         57009         5615         B         891           2176         F         431927         3896         B         56693         5616         B         890           2177         F         430046         3897         B         58586         5617	2165	F	424107	3885	В	53619	5605	В	886335
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2173         F         428558         3893         B         56326         5613         B         890           2174         F         431378         3894         B         55107         5614         B         889           2175         F         429417         3895         B         57009         5615         B         891           2176         F         431927         3896         B         56693         5616         B         890           2177         F         430046         3897         B         58586         5617         B         892           2178         F         432609         3898         B         57489         5618         B         891           2180         F         433005         3900         B         58749         5620         B         892           2181         F         431082         3901         B         60649         5621         B         892           2182         F         433712         3902         B         60086         5622         B         892           2183         F         431812         3903         B         62002         5623	2171	F	426717	3891	В	55816	5611	В	890292
2174         F         431378         3894         B         55107         5614         B         889           2175         F         429417         3895         B         57009         5615         B         891           2176         F         431927         3896         B         56693         5616         B         890           2177         F         430046         3897         B         58586         5617         B         892           2178         F         432609         3898         B         57489         5618         B         891           2180         F         433005         3900         B         58749         5619         B         892           2181         F         431082         3901         B         60649         5621         B         892           2182         F         433712         3902         B         60086         5622         B         892           2183         F         431812         3903         B         62002         5623         B         894           2184         F         436400         3905         B         64275         5624	2172	F	430475	3892	В	54416	5612	В	888879
2175         F         429417         3895         B         57009         5615         B         891           2176         F         431927         3896         B         56693         5616         B         890           2177         F         430046         3897         B         58586         5617         B         892           2178         F         432609         3898         B         57489         5618         B         891           2179         F         430710         3899         B         59394         5619         B         893           2180         F         433005         3900         B         58749         5620         B         892           2181         F         431082         3901         B         60649         5621         B         892           2182         F         433712         3902         B         60086         5622         B         892           2183         F         431812         3903         B         62002         5623         B         892           2184         F         436521         3904         B         62375         5624	2173	F	428558	3893	В	56326	5613	В	890775
2176         F         431927         3896         B         56693         5616         B         890           2177         F         430046         3897         B         58586         5617         B         892           2178         F         432609         3898         B         57489         5618         B         891           2179         F         430710         3899         B         59394         5619         B         893           2180         F         433005         3900         B         58749         5620         B         892           2181         F         431082         3901         B         60649         5621         B         892           2182         F         433712         3902         B         60086         5622         B         892           2183         F         431812         3903         B         62002         5623         B         892           2184         F         436521         3904         B         62375         5624         B         892           2185         F         436897         3906         B         61715         5626	2174	F	431378	3894	В	55107	5614	В	889595
2177         F         430046         3897         B         58586         5617         B         892           2178         F         432609         3898         B         57489         5618         B         891           2179         F         430710         3899         B         59394         5619         B         893           2180         F         433005         3900         B         58749         5620         B         892           2181         F         431082         3901         B         60649         5621         B         892           2182         F         433712         3902         B         60086         5622         B         892           2183         F         431812         3903         B         62002         5623         B         892           2184         F         436521         3904         B         62375         5624         B         892           2185         F         43640         3905         B         64275         5625         B         894           2187         F         435057         3907         B         63633         5627<	2175	F	429417	3895	В	57009	5615	В	891481
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2180         F         433005         3900         B         58749         5620         B         892           2181         F         431082         3901         B         60649         5621         B         892           2182         F         433712         3902         B         60086         5622         B         892           2183         F         431812         3903         B         62002         5623         B         894           2184         F         436521         3904         B         62375         5624         B         892           2185         F         434640         3905         B         64275         5625         B         894           2186         F         436897         3906         B         61715         5626         B         892           2187         F         435057         3907         B         63633         5627         B         892           2188         F         437882         3909         B         64601         5629         B         892           2190         F         438296         3910         B         63981         5630	2178	F	432609	3898	В	57489	5618	В	891428
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2184         F         436521         3904         B         62375         5624         B         892           2185         F         434640         3905         B         64275         5625         B         892           2186         F         436897         3906         B         61715         5626         B         893           2187         F         435057         3907         B         63633         5627         B         893           2188         F         439741         3908         B         62699         5628         B         893           2189         F         437882         3909         B         64601         5629         B         893           2190         F         438296         3910         B         63981         5630         B         893           2191         F         436377         3911         B         65858         5631         B         893           2192         F         440475         3912         B         64268         5632         B         893           2193         F         438538         3913         B         66227         5633	2182	F	433712	3902	В	60086	5622	2 B	892259
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2187         F         435057         3907         B         63633         5627         B         895           2188         F         439741         3908         B         62699         5628         B         895           2189         F         437882         3909         B         64601         5629         B         895           2190         F         438296         3910         B         63981         5630         B         895           2191         F         436377         3911         B         65858         5631         B         895           2192         F         440475         3912         B         64268         5632         B         895           2193         F         438538         3913         B         66227         5633         B         895           2194         F         440281         3914         B         64423         5634         B         895           2195         F         438394         3915         B         66309         5635         B         895	2185	F	434640	3905	В	64275	5625	5 B	894611
2188         F         439741         3908         B         62699         5628         B         893           2189         F         437882         3909         B         64601         5629         B         893           2190         F         438296         3910         B         63981         5630         B         893           2191         F         436377         3911         B         65858         5631         B         893           2192         F         440475         3912         B         64268         5632         B         893           2193         F         438538         3913         B         66227         5633         B         893           2194         F         440281         3914         B         64423         5634         B         893           2195         F         438394         3915         B         66309         5635         B         893	2186	F	436897	3906	В	61715	5626	5 B	893194
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2191     F     436377     3911     B     65858     5631     B     899       2192     F     440475     3912     B     64268     5632     B     899       2193     F     438538     3913     B     66227     5633     B     899       2194     F     440281     3914     B     64423     5634     B     899       2195     F     438394     3915     B     66309     5635     B     899	2189	F	437882	3909	В	64601	5629	9 B	895263
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2197 F 439080 3917 B 66756 5637 B 89	2197	F	439080	3917	В	66756	563	7 B	899694
2198 F 442121 3918 B 65705 5638 B 89	2198	F	442121	3918	В	65705	563	8 B	899665
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2204	F	443285	3924	В	67961	5644	В	905307
2205	F	441384	3925	В	69841	5645	В	907291
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2207	F	442406	3927	В	70662	5647	В	909083
2208	F	444472	3928	В	70984	5648	В	908055
2209	F	442568	3929	В	72885	5649	В	909955
2210	F	444960	3930	В	69392	5650	В	908358
2211	F	443040	3931	В	71314	5651	В	910273
2212	F	445556	3932	В	71365	5652	В	908900
2213	F	443681	3933	В	73287	5653	В	910831
2214	F	447565	3934	В	72253	5654	В	909607
2215	F	445676	3935	В	74167	5655	В	911450
2216	F	448396	3936	В	73916	5656	В	911760
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2218	F	450057	3938	В	76398	5658	В	912584
2219	F	448133	3939	В	78328	5659	В	914529
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2222	F	450988	3942	В	78592	5662	В	914208
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2225	F	450329	3945	В	81476	5665	В	917272
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2227	F	451581	3947	В	81861	5667	В	917747
2228	F	454643	3948	B	80203	5668	В	916886
2229	F	452718	3949	В	82108	5669	В	918778
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2231	F	454124	3951	В	82565	5671	В	919827
2232	F	456785	3952	В	81257	5672	В	919070
2233	F	454897	3953	В	83184	5673	В	920972
2234	F	457749	3954	В	83370	5674	В	920107
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2242	F	460133	3962	В	85648	5682	В	922661
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2245	F	459327	3965	В	88052	5685	В	925893
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2277	F	475181	3997	В	104947	5717	В	940928
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2283	F	477158	4003	В	106618	5723	В	942925
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2286	F	481237	4006	В	112299	5726	В	941586
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2292	F	483976	4012	В	114352	5732	В	943404
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2327	F	495210	4047	В	140361	5767	В	959202
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2352	F	513523	4072	В	155395	5792	В	971481
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2657         F         672139         4377         B         290961         6097         B         1133490           2658         F         674573         4378         B         289459         6098         B         1132846           2669         F         675234         4380         B         289914         6100         B         1134070           2661         F         675337         4381         B         2991796         6101         B         1134070           2662         F         675834         4382         B         290477         6102         B         1135089           2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         293381         6104         B         1135089           2665         F         676746         4386         B         292309         6105         B         1137155           2666         F         676746         4386         B         291463         6106         B         1136186           2667         F         674888         4387							<u></u>	В	1131563
2658         F         674573         4378         B         289459         6098         B         1132846           2669         F         672674         4379         B         291358         6099         B         1134684           2660         F         675234         4380         B         289914         6100         B         1134070           2661         F         673377         4381         B         291796         6101         B         1136016           2662         F         675834         4382         B         290477         6102         B         1135089           2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         293381         6104         B         1135089           2665         F         674477         4385         B         292309         6105         B         113715           2666         F         676746         4386         B         291463         6106         B         1136186           2667         F         678884         4389         B<				4377	В	290961	6097	В	1133490
2659         F         672674         4379         B         291358         6099         B         1134684           2660         F         675234         4380         B         289914         6100         B         1134070           2661         F         675377         4381         B         291796         6101         B         1136016           2662         F         675834         4382         B         290477         6102         B         1135089           2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         290381         6104         B         1135815           2665         F         674477         4385         B         292309         6105         B         113715           2666         F         67646         4386         B         291463         6106         B         1136186           2667         F         67488         4387         B         293372         6107         B         1138084           2669         F         675834         4389         B <td></td> <td>F</td> <td></td> <td></td> <td>В</td> <td>289459</td> <td>6098</td> <td>В</td> <td>1132846</td>		F			В	289459	6098	В	1132846
2660         F         675234         4380         B         289914         6100         B         1134070           2661         F         673377         4381         B         291796         6101         B         1136016           2662         F         675834         4382         B         290477         6102         B         1135089           2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         290381         6104         B         1135815           2665         F         67646         4386         B         292309         6105         B         1137715           2666         F         67646         4386         B         293372         6106         B         1136186           2667         F         674888         B         292307         6108         B         1136084           2669         F         675834         4389         B         293372         6107         B         1138084           2670         F         678270         4390         B         293027		F	672674	4379	В	291358	6099	В	1134684
2661         F         673377         4381         B         291796         6101         B         1136016           2662         F         675834         4382         B         290477         6102         B         1135089           2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         299381         6104         B         1137937           2665         F         676447         4385         B         292309         6105         B         113715           2666         F         676746         4386         B         291463         6106         B         1136186           2667         F         674888         B         293372         6107         B         1136084           2668         F         677769         4388         B         292104         6108         B         1137365           2669         F         675834         4389         B         293027         6110         B         1140364           2671         F         676378         4391         B         29451					В		6100	В	1134070
2662         F         675834         4382         B         290477         6102         B         1135089           2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         290381         6104         B         1135815           2665         F         674477         4385         B         292309         6105         B         1137715           2666         F         676746         4386         B         291463         6106         B         1136186           2667         F         674888         4387         B         293372         6107         B         1138084           2669         F         675834         4389         B         293372         6109         B         1139255           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1140364           2672         F         679221         4392         B					В	291796	6101	В	1136016
2663         F         673906         4383         B         292423         6103         B         1137037           2664         F         676378         4384         B         290381         6104         B         1135815           2665         F         67646         4386         B         291463         6106         B         113715           2666         F         676488         4387         B         293372         6107         B         1138084           2668         F         677769         4388         B         292104         6108         B         1137365           2669         F         675834         4389         B         293999         6109         B         1139255           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1142228           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677978         4394         B </td <td></td> <td>F</td> <td></td> <td>4382</td> <td>В</td> <td>290477</td> <td>6102</td> <td>В</td> <td>1135089</td>		F		4382	В	290477	6102	В	1135089
2664         F         676378         4384         B         290381         6104         B         1135815           2665         F         674477         4385         B         292309         6105         B         1137715           2666         F         676746         4386         B         291463         6106         B         1136186           2667         F         674888         4387         B         293372         6107         B         1138084           2669         F         675834         4389         B         293999         6109         B         1137365           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1140364           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1142478           2675         F         679778         4395         B		F	673906	4383	В	292423	6103	В	1137037
2666         F         676746         4386         B         291463         6106         B         1136186           2667         F         674888         4387         B         293372         6107         B         1138084           2668         F         677769         4388         B         292104         6108         B         1137365           2669         F         675834         4389         B         293999         6109         B         1139255           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1140364           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677925         4393         B         295409         6113         B         1142478           2675         F         677978         4395         B         295838         6115         B         1142478           2676         F         681173         4396         B		F	676378	4384	В	290381	6104	В	1135815
2667         F         674888         4387         B         293372         6107         B         1138084           2668         F         677769         4388         B         292104         6108         B         1137365           2669         F         675834         4389         B         293999         6109         B         1139255           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1142228           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         114297           2677         F         679288         4397         B<	2665	F	674477	4385	В	292309	6105	В	1137715
2668         F         677769         4388         B         292104         6108         B         1137365           2669         F         675834         4389         B         293999         6109         B         1139255           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1142228           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1147783           2677         F         679288         4397         B	2666	F	676746	4386	В	291463	6106	В	1136186
2669         F         675834         4389         B         293999         6109         B         1139255           2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1142228           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2679         F         678674         4399         B	2667	F	674888	4387	В	293372	6107	В	1138084
2670         F         678270         4390         B         293027         6110         B         1140364           2671         F         676378         4391         B         294951         6111         B         1142228           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1148953           2680         F         682210         4400         B	2668	F	677769	4388	В	292104	6108	В	1137365
2671         F         676378         4391         B         294951         6111         B         1142228           2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         682542         4402         B	2669	F	675834	4389	В	293999	6109	В	1139255
2672         F         679221         4392         B         293507         6112         B         1141611           2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         114291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2679         F         678674         4399         B         297219         6119         B         1148846           2680         F         682210         4400         B         298373         6120         B         114769           2681         F         682542         4402         B </td <td>2670</td> <td>F</td> <td>678270</td> <td>4390</td> <td>В</td> <td>293027</td> <td>6110</td> <td>В</td> <td>1140364</td>	2670	F	678270	4390	В	293027	6110	В	1140364
2673         F         677325         4393         B         295409         6113         B         1143485           2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2680         F         682210         4400         B         296373         6129         B         1147769           2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B	2671	F	676378	4391	В	294951	6111	В	1142228
2674         F         679874         4394         B         293999         6114         B         1142478           2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2679         F         678674         4399         B         297219         6119         B         1148846           2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         682542         4402         B         298114         6122         B         1148415           2683         F         682607         4403         B         299985         6123         B         1150357           2684         F         681842         4405         B	2672	F	679221	4392	В	293507	6112	В	1141611
2675         F         677978         4395         B         295838         6115         B         1144291           2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2679         F         678674         4399         B         297219         6119         B         1148846           2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         681842         4405         B	2673	F	677325	4393	В	295409	6113	В	1143485
2676         F         681173         4396         B         294889         6116         B         1145907           2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2679         F         678674         4399         B         297219         6119         B         1148846           2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         29985         6123         B         1150357           2684         F         681842         4405         B         300623         6124         B         1149762           2686         F         682410         4406         B<	2674	F	679874	4394	В	293999	6114	В	1142478
2677         F         679288         4397         B         296750         6117         B         1147783           2678         F         680607         4398         B         295312         6118         B         1146953           2679         F         678674         4399         B         297219         6119         B         1148846           2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         681842         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         682410         4406         B	2675	F	677978	4395	В	295838	6115	В	1144291
2678         F         680607         4398         B         295312         6118         B         1146953           2679         F         678674         4399         B         297219         6119         B         1148846           2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         683716         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B	2676	F	681173	4396	В	294889	6116	В	1145907
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2680         F         682210         4400         B         296373         6120         B         1147769           2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         683716         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         684312         4406         B         299027         6126         B         1149462           2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2678	F	680607	4398	В	295312	6118	В	1146953
2681         F         680303         4401         B         298305         6121         B         1149703           2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         683716         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         684312         4406         B         299027         6126         B         1149462           2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2679	F	678674	4399	В	297219	6119	В	1148846
2682         F         682542         4402         B         298114         6122         B         1148415           2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         683716         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         684312         4406         B         299027         6126         B         1149462           2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2680	F	682210	4400	В	296373	6120	В	1147769
2683         F         680607         4403         B         299985         6123         B         1150357           2684         F         683716         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         684312         4406         B         299027         6126         B         1149462           2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2681	F	680303	4401	В	298305	6121	В	1149703
2684         F         683716         4404         B         298656         6124         B         1148758           2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         684312         4406         B         299027         6126         B         1149462           2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2682	F	682542	4402	В	298114	6122	В	1148415
2685         F         681842         4405         B         300623         6125         B         1150658           2686         F         684312         4406         B         299027         6126         B         1149462           2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2683	F	680607	4403	В	299985	6123	В	1150357
2686     F     684312     4406     B     299027     6126     B     1149462       2687     F     682410     4407     B     300899     6127     B     1151258       2688     F     684880     4408     B     299805     6128     B     1149932	2684	F	683716	4404	В	298656	6124	В	1148758
2687         F         682410         4407         B         300899         6127         B         1151258           2688         F         684880         4408         B         299805         6128         B         1149932	2685	F	681842	4405	В	300623	6125	В	1150658
2688 F 684880 4408 B 299805 6128 B 1149932	2686	F	684312	4406	В	299027	6126	В	1149462
	2687	F	682410	4407	В	300899	6127	В	1151258
2689 F 682916 4409 B 301692 6129 B 1151845	2688	F	684880	4408	В	299805	6128	В	1149932
1	2689	F	682916	4409	В	301692	6129	В	1151845

2690	F	685958	4410	В	300722	6130	В	1150814
2691	F	684143	4411	В	302621	6131	В	1152747
2692	F	687264	4412	В	301846	6132	В	1151409
2693	F	685363	4413	В	303706	6133	В	1153285
2694	F	687959	4414	В	302660	6134	В	1152540
2695	F	685958	4415	В	304642	6135	В	1154341
2696	F	688514	4416	В	303066	6136	В	1154863
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2698	F	689372	4418	В	303626	6138	В	1155886
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2703	F	689372	4423	В	307390	6143	В	1159947
2704	F	692436	4424	В	306459	6144	В	1160998
2705	F	690546	4425	В	308393	6145	В	1162864
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2725	F	704105	4445	В	315784	6165	В	1171347
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2727	F	704685	4447	В	316804	6167	В	1171947
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2745	F	713725	4465	В	327001	6185	В	1181171
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2831	F	745071	4551	В	371988	6271	В	1218462
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2859	F	760141	4579	В	385161	6299	В	2068
2860	F	762611	4580	В	383789	6300	F	25997
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2909         F         781433         4629         B         409450         6349         F           2910         F         785138         4630         B         407922         6350         F           2911         F         783238         4631         B         409744         6351         F           2912         F         786197         4632         B         409039         6352         F           2913         F         784328         4633         B         410960         6353         F           2914         F         788274         4634         B         410673         6354         F           2915         F         786387         4635         B         412559         6355         F           2916         F         788679         4636         B         411193         6356         F           2917         F         786778         4637         B         413064         6357         F           2918         F         790090         4638         B         412049         6358         F           2920         F         791608         4640         B         413946	984362 996001 994109 999731 997877 1009782 1007891
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2911         F         783238         4631         B         409744         6351         F           2912         F         786197         4632         B         409039         6352         F           2913         F         784328         4633         B         410960         6353         F           2914         F         788274         4634         B         410673         6354         F           2915         F         786387         4635         B         412559         6355         F           2916         F         788679         4636         B         411193         6356         F           2917         F         786778         4637         B         413064         6357         F           2918         F         790090         4638         B         412049         6358         F           2919         F         788213         4639         B         413946         6359         F           2920         F         791608         4640         B         414525         6360         F           2922         F         792499         4642         B         415622	994109 999731 997877 1009782 1007891
2912         F         786197         4632         B         409039         6352         F           2913         F         784328         4633         B         410960         6353         F           2914         F         788274         4634         B         410673         6354         F           2915         F         786387         4635         B         412559         6355         F           2916         F         788679         4636         B         411193         6356         F           2917         F         786778         4637         B         413064         6357         F           2918         F         790090         4638         B         412049         6358         F           2919         F         788213         4639         B         413946         6359         F           2920         F         791608         4640         B         414525         6360         F           2921         F         789711         4641         B         416425         6361         F           2922         F         792499         4642         B         415622	999731 997877 1009782 1007891
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2921     F     789711     4641     B     416425     6361     F       2922     F     792499     4642     B     415622     6362     F       2923     F     790605     4643     B     417559     6363     F	1010540
2922     F     792499     4642     B     415622     6362     F       2923     F     790605     4643     B     417559     6363     F	1028431
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	1086215
	1084362
2924 F 793324 4644 B 416072 6364 F	1118417
2925 F 791440 4645 B 417968 6365 F	1116527
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2927 F 792185 4647 B 419259 6367 F	1167713
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2939	F	797649	4659	В	424158	6379	В	29730
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2969	F	810340	4689	В	439423	6409	В	597289

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2972	F	813355	t	4692	В	440583		6412	В	707138
2973	F	811466	ļ	4693	В	442491		6413	В	709063
2974	F	815198	Ì	4694	В	440583	ľ	6414	В	803951
2975	F	813243		4695	В	442441	T	6415	В	805790
2976	F	815798	İ	4696	В	441274		6416	В	849771
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2978	F	816879		4698	В	441459		6418	В	913917
2979	F	814940		4699	В	443353	ľ	6419	В	915796
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3003	F	826588	1	4723	В	453045	1	6443	В	1172218
3004	F	829813	1	4724	В	451482	1	6444	В	1181427
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3006	F	830824
3007	F	828906
3008	F	831936
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3011	F	831274

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TABLE 6

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790332G10#	6458	6654	A
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790352D2#	6462	6658	A
790373F2#	6463	6659	A
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790282F3#	6465	6661	A
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790293A5#	6470	6666	A
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790394F2#	6481	6677	В
790222G5#	6482	6678	В
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790394A10#	6586	6782	В
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790254G2#	6602	6798	В
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790213D3#	6619	6815	С
790394D2#	6620	6816	С
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790014A4#	6622	6818	С
790324H4#	6623	6819	С
790082B4#	6624	6820	С
L	<del></del>	<del></del>	1

790324A6#	6625	6821	С
790424A12#	6626	6822	С
790044G8#	6627	6823	С
790323C6#	6628	6824	С
790312G4#	6629	6825	С
790053C11#	6630	6826	С
890022B7#	6631	6827	С
790392A2#	6632	6828	С
890023D8#	6633	6829	С
790301F1#	6634	6830	C
790343A11#	6635	6831	C
790421A2#	6636	6832	С
790271G2#	6637	6833	С
790302G12#	6638	6834	С
790341E5#	6639	6835	С
790283B6#	6640	6836	С
790222A4#	6641	6837	С
790241B8#	6642	6838	С
790014C2#	6643	6839	С
790402C1#	6644	6840	С
790264E9#	6645	6841	С
790242G4#	6646	6842	С
790422F3#	6647	6843	С

TABLE 7

SEQ ID	or.	5'position
6452	В	29372
6453	В	30198
6454	В	31007
6455	В	31126
6456	В	32735
6457	В	32264
6458	В	32898
6459	В	33582
6460	В	33519
6461	В	- 34836_
6462	В	35795
6463	В	35548
6464	В	35825
6465	В	37239
6466	В	36761
6467	В	37045
6468	В	36761
6469	В	37958
6470	В	38636
6471	В	39813
6472	В	41140
6473	В	40575
6474	В	40526
6475	В	501495
6476	В	502410
6477	В	502586
6478	В	503233
6479	В	503749
6480	В	504488
6481	В	504206
6482	В	504310
6483	В	505455
6484	В	505877

SEQ ID	or.	5'position
6583	В	547718
6584	В	547184
6585	В	547684
6586	В	547342
6587	В	548946
6588	В	549071
6589	В	550054
6590	В	549989
6591	В	550426
6592	B	_550055
6593	В	550132
6594	В	550132
6595	В	551400
6596	В	551572
6597	В	551468
6598	В	550849
6599	В	552137
6600	В	552325
6601	В	552583
6602	В	553033
6603	В	553629
6604	В	553960
6605	В	553914
6606	В	554354
6607	В	555783
6608	В	555687
6609	В	556441
6610	В	557054
6611	В	556627
6612	В	557292
6613	В	557050
6614	В	815995
6615	В	817104

SEQ ID	or.	5'position
6714	F	519646
6715	F	520201
6716	F	520563
6717	F	521015
6718	F	521162
6719	F	521543
6720	F	521739
6721	F	522328
6722	F	522567
6723	F	522915
6724	F	523300
6725	F	523791
6726	F	523959
6727	F	524369
6728	F	524801
6729	F	525085
6730	F	525241
6731	F	525738
6732	F	526263
6733	F	526628
6734	F	526779
6735	F	527004
6736	F	527230
6737	F	527381
6738	F	527545
6739	F	527691
6740	F	527932
6741	F	527995
6742	F	528167
6743	F	528610
6744	F	529063
6745	F	529710
6746	F	531140

6485	В	506655
6486	В	506513
6487	В	507532
6488	В	507742
6489	В	508050
6490	В	507771
6491	В	509120
6492	В	509646
6493	В	510137
6494	В	510953
6495	В	511165
6496	В	511526
6497	В	511993
6498	В	513012
6499	В	512983
6500	В	512781
6501	В	514155
6502	В	515036
6503	В	515287
6504	В	516292
6505	В	516234
6506	В	516337
6507	В	517347
6508	В	517005
6509	В	516888
6510	В	516234
6511	В	517560
6512	В	517337
6513	В	518756
6514	В	518943
6515	В	519833
6516	В	520123
6517	В	520574
6518	В	520888
6519	В	522154
L		

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6616	В	817104
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6619	В	821017
6620	В	821379
6621	В	821504
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6624	В	823380
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6627	В	825288
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6629	В	825403
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6633	В	828146
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6635	В	827571
6636	В	828472
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6645	В	831468
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6647	В	832293
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6649	F	29043
6650	F	29656
L	٠	<u> </u>

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6753	F	533536
6754	F	533868
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6760	F	536504
6761	F	537013
6762	F	537710
6763	F	538047
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6770	F	540962
6771	F	541721
6772	F	542198
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6774	F	543180
6775	F	543877
6776	F	544601
6777	F	544866
6778	F	545442
6779	F	545948
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6781	F	546585
L	1	

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6523	В	523035	
6524	В	524995	
6525	В	523567	
6526	В	523477	
6527	В	523967	
6528	В	525211	
6529	В	525215	
6530	В	526133	
6531	В	525674	+
6532	В	526561	
6533	В	526697	
6534	В	526715	
6535	В	526844	
6536	В	527261	
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6538	В	528775	
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6543	В	527752	
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6549	В	531516	
6550	В	532154	
6551	В	532606	
6552	В	533407	
6553	В	533664	
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	ــــــــــــــــــــــــــــــــــــــ		l

6651	F	30157
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6653	F	31175
6654	F	31658
6655	F	31902
6656	F	32638
6657	F	33203
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6659	F	34164
6660	F	34426
6661	F	35131
6662	F	35675
6663	F	36097
6664	F	36641
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6713	F	519133
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6838	F	828729
6839	F	830099
6840	F	830281
6841	F	830491
6842	F	830550
6843	F	830576

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## WHAT IS CLAIMED IS:

	1- An isolated polynugenome, comprising	ıcleotic	le having a nucleotide sequence of a Chlamydia pneumoniae
5		(a)	the a nucleotide sequence of SEQ ID No. 1;
		(b)	the nucleotide sequence contained within the Chlamydia
		•	pneumoniae genomic DNA in ATCC Deposit No;
		(c)	the nucleotide sequence contained in a clone insert in ATCC
		` '	Deposit No;
10		(d)	a nucleotide sequence exhibiting at least 99.9% identity with the
		` '	sequence of SEQ ID No. 1; or
		(e)	a nucleotide sequence exhibiting at least 80% homology to SEQ
		( )	ID No. 1.
	- w <del>-</del>		
15			de which hybridizes to SEQ ID No. 1 or to the Chlamydia
	•		contained in ATCC deposit No or to a clone insert in
	ATCC Deposit No		under conditions of high stringency.
20	intermediate stringenc	y. cleotid	contained in ATCC deposit No under conditions of the having a nucleotide sequence of an open reading frame (ORF)
25	of a Chiamyala phean	(a)	a nucleotide sequence chosen from one of ORF2 to ORF 1297;
25		(a) (b)	a nucleotide sequence exhibiting at least 99.9% identity with
		(0)	one of ORF2 to ORF 1297; or
		(c)	a nucleotide sequence exhibiting at least 80% homology to one
		(0)	of ORF2 to ORF 1297.
30			
	5- An isolated polyconditions of high stri		ide which hybridizes to one of ORF2 to ORF 1297 under 7.
	6 An isolated make	mualaat	tide which hybridizes to one of ORF2 to ORF 1297 under
25	conditions of intermed		
33	Conditions of Intermet	uiaic si	
	7- The polynucleotide fragments thereof:	e of Cla	aims 2, 3, 4, 5, or 6 which encodes the following polypeptides or
	-	(a)	a Chlamydia pneumoniae transmembrane polypeptide having
40	•	•	between 1 and 3 transmembrane domains;

	(b)	a Chlamydia pneumoniae transmembrane polypeptide having
		between 4 and 6 transmembrane domains;
	(c)	a Chlamydia pneumoniae transmembrane polypeptide having at
		least 7 transmembrane domains;
5	(d)	a Chlamydia pneumoniae polypeptide involved in intermediate
		metabolism of sugars and/or cofactors;
	(e)	a Chlamydia pneumoniae polypeptide involved in intermediate
		metabolism of nucleotides or nucleic acids;
	<b>(f)</b>	a Chlamydia pneumoniae polypeptide involved in metabolism
10		of amino acids or polypeptides;
	(g)	a Chlamydia pneumoniae polypeptide having involved in
		metabolism of fatty acids;
	(h)	a Chlamydia pneumoniae polypeptide involved in the synthesis
		of the cell wall;
15	(i)	a Chlamydia pneumoniae polypeptide involved in transcription,
		translation, and/or maturation process;
	<b>(i)</b>	a Chlamydia pneumoniae transport polypeptide;
	(k)	a Chlamydia pneumoniae polypeptide involved in the virulence
		process;
20	(1)	a Chlamydia pneumoniae polypeptide involved in the secretory
		system and/or which is secreted;
	(m)	a Chlamydia pneumoniae polypeptide of the cellular envelope
		or outer cellular envelope of Chlamydia pneumoniae.
	(n)	a Chlamydia pneumoniae surface exposed polypeptide;
25	(o)	a Chlamydia pneumoniae lipoprotein;
	(p)	a Chlamydia pneumoniae polypeptide involved in
		lipopolysaccharide biosynthesis;
	(q)	a Chlamydia pneumoniae KDO-related polypeptide;
	(r)	a Chlamydia pneumoniae phosphomannomutase-related
30		polypeptide;
	(s)	a Chlamydia pneumoniae lipid A component-related
		polypeptide;
	(t)	a Chlamydia pneumoniae phosphoglucomutase-related
		polypeptide;
35	(u)	a Chlamydia pneumoniae polypeptide that contains an RGD
		sequence;
	(v)	a Chlamydia pneumoniae Type III secreted polypeptide;
	(w)	a Chlamydia pneumoniae cell wall anchored surface
		polypeptide; or

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- (x) a Chlamydia pneumoniae polypeptide that is not found in Chlamydia trachomatis.
- 8- A polynucleotide encoding a fusion protein, comprising one of ORF2 to ORF1297 of 5 Claim 4, 5, or 6 ligated in frame to a polynucleotide encoding a heterologous polypeptide.
  - 9- A recombinant vector that contains the polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
  - 10- A recombinant vector that contains the polynucleotide of Claim 8.

11- A recombinant vector that contains the polynucleotide of Claim 4, 5 or 6, operatively associated with a regulatory sequence that controls gene expression.

- 12- A recombinant vector that contains the polynucleotide of Claim 8 operatively associated with a regulatory sequence that controls gene expression.
  - 13- A genetically engineered host cell that contains the polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
- 20 14- A genetically engineered host cell that contains the polynucleotide of Claim 8.
  - 15- A genetically engineered host cell that contains the polynucleotide of Claim 4, 5 or 6 operatively associated with a regulatory sequence that controls gene expression in the host cell.

16- A genetically engineered host cell that contains the polynucleotide of Claim 8 operatively associated with a regulatory sequence that controls gene expression in the host cell.

17- A method for producing a polypeptide, comprising:

- (a) culturing the genetically engineered host cell of Claim 15 under conditions suitable to produce the polypeptide encoded by the polynucleotide; and
- (b) recovering the polypeptide from the culture.
- 35 18- A method for producing a fusion protein, comprising:
  - (a) culturing the genetically engineered host cell of Claim 16 under conditions suitable to produce the fusion protein encoded by the polynucleotide; and
  - (b) recovering the fusion protein from the culture.

- 19- A polypeptide encoded by the polynucleotide of Claim 4, 5 or 6.
- 20- The polypeptide of Claim 19 which immunoreacts with seropositive serum of an 5 individual infected with *Chlamydia pneumoniae*.
  - 21- The polypeptide of Claim 19 which comprises the following polypeptides or fragments thereof:
    - (a) a *Chlamydia pneumoniae* transmembrane polypeptide having between 1 and 3 transmembrane domains;
    - (b) a Chlamydia pneumoniae transmembrane polypeptide having between 4 and 6 transmembrane domains;
    - (c) a *Chlamydia pneumoniae* transmembrane polypeptide having at least 7 transmembrane domains;
    - (d) a *Chlamydia pneumoniae* polypeptide involved in intermediate metabolism of sugars and/or cofactors;
    - (e) a Chlamydia pneumoniae polypeptide involved in intermediate metabolism of nucleotides or nucleic acids;
    - (f) a Chlamydia pneumoniae polypeptide involved in metabolism of amino acids or polypeptides;
    - (g) a Chlamydia pneumoniae polypeptide involved in metabolism of fatty acids;
    - (h) a Chlamydia pneumoniae polypeptide involved in the synthesis of the cell wall;
    - (i) a Chlamydia pneumoniae polypeptide involved in transcription, translation, and/or maturation process;
    - (j) a Chlamydia pneumoniae transport polypeptide;
    - (k) a Chlamydia pneumoniae polypeptide involved in the virulence process:
    - (l) a Chlamydia pneumoniae polypeptide involved in the secretory system and/or which is secreted;
    - (m) a Chlamydia pneumoniae polypeptide of the cellular envelope or outer cellular envelope of Chlamydia pneumoniae.
    - (n) a Chlamydia pneumoniae surface exposed polypeptide;
    - (o) a Chlamydia pneumoniae lipoprotein;
    - (p) a *Chlamydia pneumoniae* polypeptide involved in lipopolysaccharide biosynthesis;
    - (q) a Chlamydia pneumoniae KDO-related polypeptide;

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phosphomannomutase-related Chlamydia pneumoniae (r) polypeptide; phosphoglucomutase-related Chlamydia (s) pneumoniae polypeptide; a Chlamydia component-related pneumoniae lipid (t) polypeptide; a Chlamydia pneumoniae polypeptide that contains an RGD (u) sequence; a Chlamydia pneumoniae Type III secreted polypeptide; (v) a Chlamydia pneumoniae cell wall anchored surface (w) polypeptide; or

a Chlamydia pneumoniae polypeptide that is not found in

15 22- A fusion protein encoded by the polynucleotide of Claim 8.

(x)

- 23- The fusion protein of Claim 22 which immunoreacts with seropositive serum of an individual infected with *Chlamydia pneumoniae*.
- 20 24- An antibody that immunospecifically binds to the polypeptide of Claim 19.
  - 25- An antibody that immunospecifically binds to the fusion protein of Claim 22.

Chlamydia trachomatis.

- 26- A method for the detection and/or identification of *Chlamydia pneumoniae* in a biological sample, comprising:
  - (a) contacting the sample with a polynucleotide primer of Claim 1,
     2, 3, 4, 5, or 6 in the presence of a polymerase enzyme and nucleotides under conditions which permit primer extension;
     and
  - (b) detecting the presence of primer extension products in the sample in which the detection of primer extension products indicates the presence of *Chlamydia pneumoniae* in the sample.
- 27- A method for the detection and/or identification of *Chlamydia pneumoniae* in a biological sample, comprising:
  - (a) contacting the sample with a polynucleotide probe of Claim 1,
    2, 3, 4, 5, or 6 under conditions which permit hybridization of complementary base pairs; and

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- (b) detecting the presence of hybridization complexes in the sample in which the detection of hybridization complexes indicates the presence of *Chlamydia pneumoniae* in the sample.
- 5 28- A method for the detection and/or identification of *Chlamydia pneumoniae* in a biological sample, comprising:
  - (a) contacting the sample with the antibody of Claim 24 under conditions suitable for the formation of immune complexes; and
  - (b) detecting the presence of immune complexes in the sample, in which the detection of immune complexes indicates the presence of *Chlamydia pneumoniae* in the sample.
  - 29- A method for the detection and/or identification of antibodies to *Chlamydia pneumoniae* in a biological sample, comprising:
    - (a) contacting the sample with a polypeptide of Claim 19 under conditions suitable for the formation of immune complexes; and
    - (b) detecting the presence of immune complexes in the sample, in which the detection of immune complexes indicates the presence of *Chlamydia pneumoniae* in the sample.
  - 30- A DNA chip containing an array of polynucleotides comprising at least one of the polynucleotides of Claim 1, 2, 3, 4, 5, or 6.
- 31- A protein chip containing an array of polypeptides comprising at least one of the polypeptides of Claim 19.
  - 32- An immunogenic composition comprising the polypeptide of Claim 19 and a pharmaceutically acceptable carrier.
- 30 33- An immunogeneic composition comprising the polypeptide of Claim 20 and a pharmaceutically acceptable carrier.
  - 34- An immunogenic composition comprising the fusion protein of Claim 22 and a pharmaceutically acceptable carrier.
  - 35- An immunogenic composition comprising the fusion protein of Claim 23 and a pharmaceutically acceptable carrier.

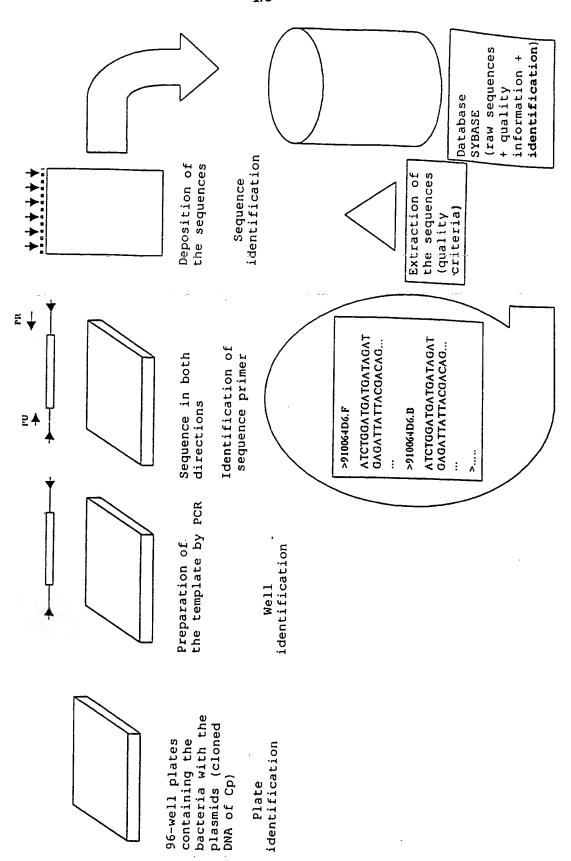
- 36- A pharmaceutical composition comprising the polypeptide of Claim 19 and a pharmaceutically acceptable carrier.
- 37- A pharmaceutical composition comprising the polypeptide of Claim 20 and a pharmaceutically acceptable carrier.
  - 38- A pharmaceutical composition comprising the polypeptide of Claim 22 and a pharmaceutically acceptable carrier.
- 10 39- A pharmaceutical composition comprising the polypeptide of Claim 23 and a pharmaceutically acceptable carrier.
  - 40- A method of immunizing against *Chlamydia pneumoniae*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 32.
  - 41- A method of immunizing against *Chlamydia pneumoniae*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 33.
- 42- A method of immunizing against *Chlamydia pneumoniae*, comprising administering to a host an immunizing amount of the immunogenic composition of Claim 34.
  - 43- A method of immunizing against *Chlamydia pneumoniae*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 35.
- 25 44- A DNA immunogenic composition comprising the expression vector of Claim 11.
  - 45- The DNA composition of Claim 44, wherein the DNA composition directs the expression of a neutralizing epitope of *Chlamydia pneumoniae*.
- 30 46- A DNA immunogenic composition comprising the expression vector of Claim 12.
  - 47- The DNA composition of Claim 46, wherein the DNA composition directs the expression of a neutralizing epitope of *Chlamydia pneumoniae*.
- 35 48- A screening assay, comprising:
  - (a) contacting a test compound with an isolated polynucleotide of Claim 1, 2, 3, 4, 5 or 6; and
  - (b) detecting whether binding occurs.

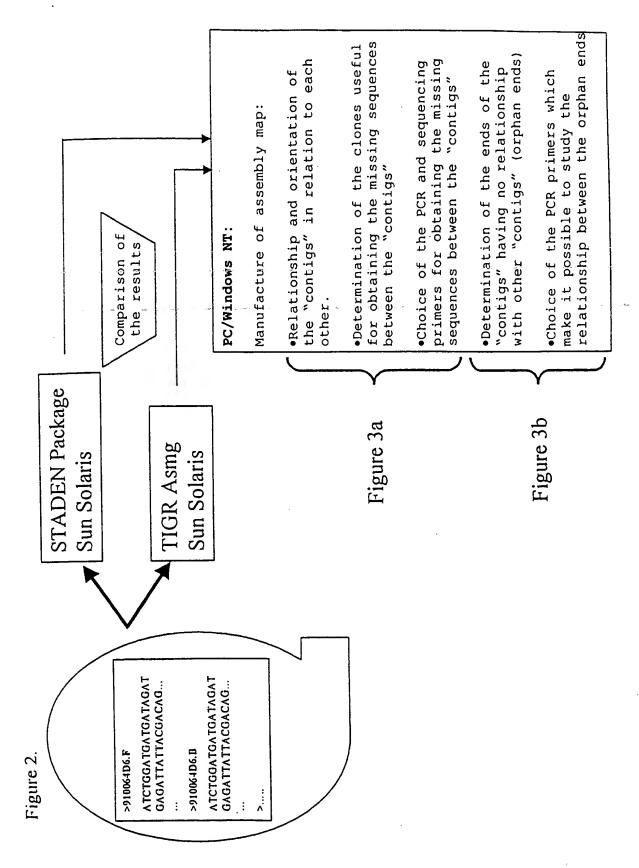
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- 49- A screening assay, comprising:
  - (a) contacting a test compound with the polypeptide of Claim19; and
  - (b) detecting whether binding occurs.
- 50- A screening assay, comprising:
  - (a) contacting a test compound with the polypeptide of Claim 22; and
  - (b) detecting whether binding occurs.
- 10 51- A kit comprising a container containing an isolated polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
  - 52- The kit of Claim 51 wherein the polynucleotide is a primer or a probe.
- 15 53- The kit of Claim 51 wherein the polynucleotide is a primer and the kit further comprises a container containing a polymerase.
  - 54- The kit of Claim 51 which further comprises a container containing deoxynucleotide triphosphates.
  - 55- A kit comprising a container containing an antibody that immunospecifically binds to the polypeptide of Claim 19.
- 56- A kit comprising a container containing an antibody that immunospecifically binds to the fusion protein of Claim 22.

Figure 1.





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PIGURE 3A

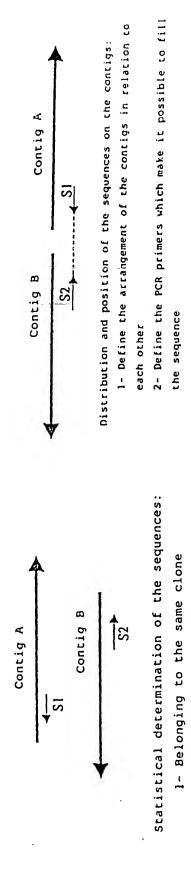
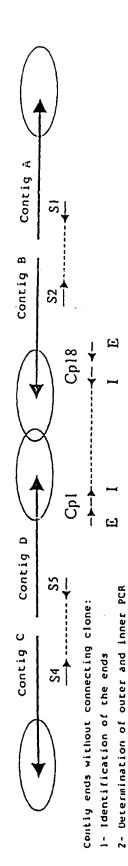


FIGURE 3B



E: outer primers

between the contigs

primers for studying the relationships

I: inner primers

SUBSTITUTE SHEET (RULE 26)

2- Situated on two different contigs

## SEQUENCE LISTING

<110>Genset SA <120>Chlamydia pneumoniae genomic sequence and polypeptides, fragments thereof and uses thereof, in particular for the diagnosis, prevention and treatment of infection <151>1997-11-21 <160>6849 <210>1 <211>1230025 <212>DNA <213>Chlamydia pneumoniae atagaaaact attaaaaaat cattgattct gtcgggaaag tatgcggata aaattcagag 60 agaataagga gaggaagatg acaaggcaga gttatgtttt gggcaattgg aaaatgcaca 120 aaacaatcca agaagctaaa gagtatgttc aaacattagc ttctntacta caaggagaac 180 ctctttcctg cactataggc atagcttctc catttacctc tttgagagcg attcatgaga 240 tgataaacac tacgggagct tttctctggt tgggagcaca aaatgtccat cccgagcttt 300 cgggtgcttt tactggagaa atttccttac ctatgcttaa ggaggtagga gtggaatttg 360 ttttagtagg tcactccgag cgtcgtcata tttttggaga gagtgatgcc tttattgctt 420 480 caaaggtaaa gtotgtagot caggogggac togtgootgt totttgtgtt ggagagagot tagaagttcg tgaagaggga aaggcgcatc aggtaatcaa aaaacagttg cttttgggat 540 tggaacagat ggataatggt tccgaatttt tgatcgccta tgaaccagta tgggctatcg 600 gcacagggaa ggtggcagaa gcttcggatg tgcaagatat tcatatgttt tgtcgtgagg 660 tagtggcaga gaggttctca gaagctacag ctgaagagat ttcgattttg tacggaggat 720 ctgtgaaggt cgataatgct cagcgatttg ggcaatgtag cgacgtcgat ggtcttttag 780 ttggcggant tctttagang ggcaaagttt ttttgaagtc gctaaaaatt ttaatgtata 840 900 atttgtgaga gttatgagat ttttttgtct attttttctt gggttcctag gatcttttca 960 ttgtgttgct gaagacaagg gcgtggattt atttggagtc tgggacgata accaaattac 1020 agagtgtgac gatagttaca tgacagaggg tcgtgaagag gttgaaaagg tagtggacgc ttagtccatc ggcttttatt tatattctcc ctaaggaagt cctgtattga agatcgcttt 1080 ctcatagata gaagtaattt tcagatagtc aataattggt ttttttaaga gaatgctagg 1140 caqqtqctcg tgtttgggca tttgattaag tctacatgaa tctggaggga gagattcttc 1200 tggtattgag aagtagaaca aaaaacaagg atcagacgtt ctccgatgtc ttcctaatcg 1260 atgtctttaa ataaggagat tggcatgaca gtgttgtttt acgcattttt attcattttc 1320 ctttttctat gtgtaattct ttgtggctta atcctggttc aagagagtaa gagcatgggg 1380 1440 ttaggttctt cgttcggcgt ggattctgga gattctgtct ttggtgtctc tactccagat 1500 attttgaaaa aagtgacttc atngtgtgct gttgctttct gcataggttg tttactactt tcattttcca cqaatctctt ggggaaaaag ttagatgcta aagaatttct attgcctgct 1560 gctgaggaga gcgacactca agcttcttct gagagcgttg aagcagatga atcctagcct 1620 atttgcggaa ttaggtgttg tctagattga agtgcaataa agctagcaag tttttatctt 1680 catacgagat atgagtgtac ggtcggataa gagtagaaat ctttcttttg ttcctatggt 1740 taagaagtcc tttggcttcc ttaaagagta tgactcttat caacccaaga aatgttttag 1800 atccaaqtqc ttqtcqtacq agatttcttc acagagctct gccaagccca tgttcagact 1860 atgattagac gtttagaata ttacggcagt cctattttaa ggaaaaagtc ttccccaatt 1920 qcaqaqatca cagatgagat tcgtaatctc gtgagtgata tgtgtgatac tatggaagca 1980 catcgtggtq tcggtttagc cgctcctcag gtagggaaaa acgtcagttt atttgtcatg 2040 2100 tgtgtagata gagagactga ggatggagag ttgattttct ctgagtctcc gagggtattt 2160 atcaatcctg ttctatcaga tccttctgaa accccgatca taggtaaaga aggatgtctt tctattcctg gattgcgagg agaagtattc cgccctcaga aaatcacagt gaccgctatg 2220 gatctcaatg gtaaaatatt tactgagcac ttggaaggat tcactgcacg tatcattatg 2280 cacgagactg accatctgaa tggagttctc tatattgatc ttatggaaga acccaaagat 2340 2400 cctaaaaaat ttaaagcctc tttagagaag atcaaacgtc gctacaatac acacttgagt aaaqaaqaac taqtttctta attgctcttc agtctgatgt aggtgatatt ttcttgtctc 2460 ttqcqtcaca tttqttqtca gctttgctta tttccccgaa caaatttcgt caaaggtttt 2520 aaaatgtgtc ttgctgattt ttgctaagag ctctttccct cgttgcttag cgatctctct 2580 tcctgctgct ttgacattga atccagcacc tttaggaagc tgtacttgat attgttcttc 2640 caacttctgt atcgactgta caaatgcatc tctagccaat atagaagctg ctgctacgac 2700 tacatcttgt tctgcacgtg gcttttgtat taaagtaata tcggtttctt ttttttgaag 2760 tgctttgagt agggtgtatt ctgaagctgc aaactgatct gaaatagcaa agacatctcc 2820 tgcaggtttg ggtgctaagt tgttgataac agtagcgtgg gcccaagcaa gaagtgtatt 2880 taaattctgg aatttcccat atagctcgtt atatttttct gggtatagaa tgatgacatc 2940 gcagacacat agtgagcgta tgatacgtgc taaagaagcg attttcgtgt ctttgagatt 3000 tttagagtct tggactttat tctcatagag tttttttaag atctctgcat tcgatgcata 3060 3120 gactgccgca atacataaag ggccaaaaaa atcacettte cetgatteat egacteccaa

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